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(54) Title: 87 HUMAN SECRETED PROTEINS

(57) Abstract

The present invention relates to 87 novel human secreted proteins and isolated nucleic acids containing the coding regions of the genes encoding such proteins. Also provided are vectors, host cells, antibodies, and recombinant methods for producing human secreted proteins. The invention further relates to diagnostic and therapeutic methods useful for diagnosing and treating disorders related to these novel human secreted proteins.

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87 Human Secreted Proteins

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Field of the Invention

This invention relates to newly identified polynucleotides and the polypeptides encoded by these polynucleotides, uses of such polynucleotides and polypeptides, and their production.

Background of the Invention

Unlike bacterium, which exist as a single compartment surrounded by a membrane, human cells and other eucaryotes are subdivided by membranes into many functionally distinct compartments. Each membrane-bounded compartment, or organelle, contains different proteins essential for the function of the organelle. The cell uses "sorting signals," which are amino acid motifs located within the protein, to target proteins to particular cellular organelles.

One type of sorting signal, called a signal sequence, a signal peptide, or a leader sequence, directs a class of proteins to an organelle called the endoplasmic reticulum (ER). The ER separates the membrane-bounded proteins from all other types of proteins. Once localized to the ER, both groups of proteins can be further directed to another organelle called the Golgi apparatus. Here, the Golgi distributes the proteins to vesicles, including secretory vesicles, the cell membrane, lysosomes, and the other organelles.

Proteins targeted to the ER by a signal sequence can be released into the extracellular space as a secreted protein. For example, vesicles containing secreted proteins can fuse with the cell membrane and release their contents into the extracellular space - a process called exocytosis. Exocytosis can occur constitutively or after receipt of a triggering signal. In the latter case, the proteins are stored in secretory vesicles (or secretory granules) until exocytosis is triggered. Similarly, proteins residing on the cell membrane can also be secreted into the extracellular space by proteolytic cleavage of a "linker" holding the protein to the membrane.

Despite the great progress made in recent years, only a small number of genes encoding human secreted proteins have been identified. These secreted proteins include the commercially valuable human insulin, interferon, Factor VIII, human growth hormone, tissue plasminogen activator, and erythropoeitin. Thus, in light of the pervasive role of secreted proteins in human physiology, a need exists for identifying and characterizing novel human secreted proteins and the genes that encode them. This knowledge will allow one to detect, to treat, and to prevent medical disorders by using secreted proteins or the genes that encode them.

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Summary of the Invention

The present invention relates to novel polynucleotides and the encoded polypeptides. Moreover, the present invention relates to vectors, host cells, antibodies, and recombinant methods for producing the polypeptides and polynucleotides. Also provided are diagnostic methods for detecting disorders related to the polypeptides, and therapeutic methods for treating such disorders. The invention further relates to screening methods for identifying binding partners of the polypeptides.

Detailed Description

Definitions

The following definitions are provided to facilitate understanding of certain terms used throughout this specification.

In the present invention, "isolated" refers to material removed from its original environment (e.g., the natural environment if it is naturally occurring), and thus is altered "by the hand of man" from its natural state. For example, an isolated polynucleotide could be part of a vector or a composition of matter, or could be contained within a cell, and still be "isolated" because that vector, composition of matter, or particular cell is not the original environment of the polynucleotide.

In the present invention, a "secreted" protein refers to those proteins capable of being directed to the ER, secretory vesicles, or the extracellular space as a result of a signal sequence, as well as those proteins released into the extracellular space without necessarily containing a signal sequence. If the secreted protein is released into the extracellular space, the secreted protein can undergo extracellular processing to produce a "mature" protein. Release into the extracellular space can occur by many mechanisms, including exocytosis and proteolytic cleavage.

As used herein, a "polynucleotide" refers to a molecule having a nucleic acid sequence contained in SEQ ID NO:X or the cDNA contained within the clone deposited with the ATCC. For example, the polynucleotide can contain the nucleotide sequence of the full length cDNA sequence, including the 5' and 3' untranslated sequences, the coding region, with or without the signal sequence, the secreted protein coding region, as well as fragments, epitopes, domains, and variants of the nucleic acid sequence. Moreover, as used herein, a "polypeptide" refers to a molecule having the translated amino acid sequence generated from the polynucleotide as broadly defined.

In the present invention, the full length sequence identified as SEQ ID NO:X was often generated by overlapping sequences contained in multiple clones (contig

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analysis). A representative clone containing all or most of the sequence for SEQ ID NO:X was deposited with the American Type Culture Collection ("ATCC"). As shown in Table 1, each clone is identified by a cDNA Clone ID (Identifier) and the ATCC Deposit Number. The ATCC is located at 10801 University Boulevard, Manassas, Virginia 20110-2209, USA. The ATCC deposit was made pursuant to the terms of the Budapest Treaty on the international recognition of the deposit of microorganisms for purposes of patent procedure.

A "polynucleotide" of the present invention also includes those polynucleotides capable of hybridizing, under stringent hybridization conditions, to sequences contained in SEQ ID NO:X, the complement thereof, or the cDNA within the clone deposited with the ATCC. "Stringent hybridization conditions" refers to an overnight incubation at 42° C in a solution comprising 50% formamide, 5x SSC (750 mM NaCl, 75 mM sodium citrate), 50 mM sodium phosphate (pH 7.6), 5x Denhardt's solution, 10% dextran sulfate, and $20 \,\mu\text{g/ml}$ denatured, sheared salmon sperm DNA, followed by washing the filters in 0.1x SSC at about 65° C.

Also contemplated are nucleic acid molecules that hybridize to the polynucleotides of the present invention at lower stringency hybridization conditions. Changes in the stringency of hybridization and signal detection are primarily accomplished through the manipulation of formamide concentration (lower percentages of formamide result in lowered stringency); salt conditions, or temperature. For example, lower stringency conditions include an overnight incubation at 37°C in a solution comprising 6X SSPE (20X SSPE = 3M NaCl; 0.2M NaH₂PO₄; 0.02M EDTA, pH 7.4), 0.5% SDS, 30% formamide, 100 ug/ml salmon sperm blocking DNA; followed by washes at 50°C with 1XSSPE, 0.1% SDS. In addition, to achieve even lower stringency, washes performed following stringent hybridization can be done at higher salt concentrations (e.g. 5X SSC).

Note that variations in the above conditions may be accomplished through the inclusion and/or substitution of alternate blocking reagents used to suppress background in hybridization experiments. Typical blocking reagents include Denhardt's reagent, BLOTTO, heparin, denatured salmon sperm DNA, and commercially available proprietary formulations. The inclusion of specific blocking reagents may require modification of the hybridization conditions described above, due to problems with compatibility.

Of course, a polynucleotide which hybridizes only to polyA+ sequences (such as any 3' terminal polyA+ tract of a cDNA shown in the sequence listing), or to a

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complementary stretch of T (or U) residues, would not be included in the definition of "polynucleotide," since such a polynucleotide would hybridize to any nucleic acid molecule containing a poly (A) stretch or the complement thereof (e.g., practically any double-stranded cDNA clone).

The polynucleotide of the present invention can be composed of any polyribonucleotide or polydeoxribonucleotide, which may be unmodified RNA or DNA or modified RNA or DNA. For example, polynucleotides can be composed of single-and double-stranded DNA, DNA that is a mixture of single- and double-stranded regions, single- and double-stranded RNA, and RNA that is mixture of single- and double-stranded regions, hybrid molecules comprising DNA and RNA that may be single-stranded or, more typically, double-stranded or a mixture of single- and double-stranded regions. In addition, the polynucleotide can be composed of triple-stranded regions comprising RNA or DNA or both RNA and DNA. A polynucleotide may also contain one or more modified bases or DNA or RNA backbones modified for stability or for other reasons. "Modified" bases include, for example, tritylated bases and unusual bases such as inosine. A variety of modifications can be made to DNA and RNA; thus, "polynucleotide" embraces chemically, enzymatically, or metabolically modified forms.

The polypeptide of the present invention can be composed of amino acids joined to each other by peptide bonds or modified peptide bonds, i.e., peptide isosteres, and may contain amino acids other than the 20 gene-encoded amino acids. The polypeptides may be modified by either natural processes, such as posttranslational processing, or by chemical modification techniques which are well known in the art. Such modifications are well described in basic texts and in more detailed monographs, as well as in a voluminous research literature. Modifications can occur anywhere in a polypeptide, including the peptide backbone, the amino acid side-chains and the amino or carboxyl termini. It will be appreciated that the same type of modification may be present in the same or varying degrees at several sites in a given polypeptide. Also, a given polypeptide may contain many types of modifications. Polypeptides may be branched, for example, as a result of ubiquitination, and they may be cyclic, with or without branching. Cyclic, branched, and branched cyclic polypeptides may result from posttranslation natural processes or may be made by synthetic methods. Modifications include acetylation, acylation, ADP-ribosylation, amidation, covalent attachment of flavin, covalent attachment of a heme moiety, covalent attachment of a nucleotide or nucleotide derivative, covalent attachment of a lipid or lipid derivative, covalent attachment of phosphotidylinositol, cross-linking, cyclization, disulfide bond formation, demethylation, formation of covalent cross-links, formation of cysteine,

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formation of pyroglutamate, formylation, gamma-carboxylation, glycosylation, GPI anchor formation, hydroxylation, iodination, methylation, myristoylation, oxidation, pegylation, proteolytic processing, phosphorylation, prenylation, racemization, selenoylation, sulfation, transfer-RNA mediated addition of amino acids to proteins such as arginylation, and ubiquitination. (See, for instance, PROTEINS -STRUCTURE AND MOLECULAR PROPERTIES, 2nd Ed., T. E. Creighton, W. H. Freeman and Company, New York (1993); POSTTRANSLATIONAL COVALENT MODIFICATION OF PROTEINS, B. C. Johnson, Ed., Academic Press, New York, pgs. 1-12 (1983); Seifter et al., Meth Enzymol 182:626-646 (1990); Rattan et al., Ann NY Acad Sci 663:48-62 (1992).)

"SEQ ID NO:X" refers to a polynucleotide sequence while "SEQ ID NO:Y" refers to a polypeptide sequence, both sequences identified by an integer specified in Table 1.

"A polypeptide having biological activity" refers to polypeptides exhibiting activity similar, but not necessarily identical to, an activity of a polypeptide of the present invention, including mature forms, as measured in a particular biological assay, with or without dose dependency. In the case where dose dependency does exist, it need not be identical to that of the polypeptide, but rather substantially similar to the dose-dependence in a given activity as compared to the polypeptide of the present invention (i.e., the candidate polypeptide will exhibit greater activity or not more than about 25-fold less and, preferably, not more than about tenfold less activity, and most preferably, not more than about three-fold less activity relative to the polypeptide of the present invention.)

Polynucleotides and Polypeptides of the Invention

FEATURES OF PROTEIN ENCODED BY GENE NO: 1

The translation product of this gene shares sequence homology with nucleolin, which is thought to be important in macromolecule binding, as well as some membrane proteins. Preferred polypeptide fragments comprise the amino acid sequence: DPEAADSGEPQNKRTPDLPEEEYVKEEIQENEEAVKKMLVEATREFEEVVVDES (SEQ ID NO:239); QKLKRKAEEDPEAADSGEPQNKRTPDLPEEEYVKEEIQENEE AVKKMLVEATREFEEVVVDES (SEQ ID NO:240); KAMEKSSLTQHSWQSLKDR YLKHLRGQEHKYLLGDAPVSPSSQKLKRKAEEDPEAADSGEPQNKRTPDLPEE EYVKEEIQENEEAVKKMLVEATREFEEVVVDESPPDFEIHI (SEQ ID NO:241). 35 Also preferred are the polynucleotide fragments encoding these polypeptide fragments.

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This gene maps to chromosome 16, and therefore can be used as a marker in linkage analysis for chromosome 16.

This gene is expressed primarily in brain and kidney and to a lesser extent in wide range of tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cell-cell interaction or cell-matrix interaction. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the brain and kidney, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., brain and other tissue of the nervous system, and kidney, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:125 as residues: Met-1 to Trp-10.

The tissue distribution and homology to nucleolin indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment/diagnosis of diseases involving cell-cell interaction or cell-extracellular matrix interaction.

FEATURES OF PROTEIN ENCODED BY GENE NO: 2

The translation product of this gene shares sequence homology with a porcine zona pellucida protein ZPDS.1711. (See Accession No. R39356.) These two proteins have weak homology with *Drosophila* commissureless and metal homeostasis proteins which are thought to be important in controlling growth cone guidance across the CNS midline and protecting cells against reactive oxygen toxicity. thus, based on homology, it is likely that this gene also be involved in development. Preferred polypeptide fragments comprise the amino acid sequence: LPSYDEAERTKAEATIPLVPGRDEDF VGRDDFDDADQLRIGNDGIFMLTFFMAFLFNWIGFFLSFCLTTSAAGRYGAISG FGLSLIKWILIVRFSTYFPGYFDGQYWLWWVFLVLGFLLFLRGFINYAKVRKM PETFSNLPRTRVLFI (SEQ ID NO:242); and/or AGRYGAISGFGLSLIKWILIVRFS (SEQ ID NO:243). Also preferred are polynucleotide fragments encoding these polypeptide fragments. This gene maps to chromosome 5, and therefore can be used in linkage analysis as a marker for chromosome 5.

This gene is expressed primarily in kidney, adrenal gland, brain and to a lesser extent in wide range of tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, fertilization control or tissue damages by metabolites or other toxic agents. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the reproductive and urosecretion system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., kidney, adrenal gland, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to zona pellucida protein indicates that polynucleotides and polypeptides corresponding to this gene are useful for fertility control such as controceptive development. The homology with metal homeostasis and commissureless genes indicates the gene's function in spermatozoa guidance and protection. It would also be useful for the treatment/diagnosis of tissue damages caused by toxic metabolites and other agents since the gene product is also expressed in urosecretive tissues.

25 FEATURES OF PROTEIN ENCODED BY GENE NO: 3

This gene is expressed primarily in liver and to a lesser extent in placenta. Preferred polypeptide fragments comprise the amino acid sequence: MKHLSAWNFT KLTFLQLWEI FEGSVENCQTLTSYSKLQIKYTFSRGSTFYI (SEQ ID NO:244). Also preferred are polynucleotide fragments encoding these polypeptide fragments.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, digestive and nutrient transport/utilization disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the digestive and

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circulatory system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., liver, and placenta, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution in liver and placenta indicates that the protein product is either an extracellular enzyme or a molecule carrier. Therefore, polynucleotides and polypeptides corresponding to this gene are useful for diagnosis/treatment of digestive and nutrient transport/utilization disorders, including malabsorption and malnutrition.

FEATURES OF PROTEIN ENCODED BY GENE NO: 4

This gene shares homology with the sap47 gene of Drosophila melanogaster, a gene which codes for a conserved neuronal protein associated with synaptic terminals. (See Mol. Brain Res. 32:45-54 (1995); see also, Accession No. 929571.) Thus, based on 15 homology, the gene of the present invention also should be associated with synaptic terminals. Preferred polypeptide fragments comprise the amino acid sequence: FSSDFRTSPWESRRVESKATSARCGLWGSGPRRRPASGMFRGLSSWLGLQQP VAGGGQPNGDAPPEQPSETVAESAEEELQQAGDQELLHQAKDFGNYLFNFASA ATKKITESVAETAQTIKKSVEEGKIDGIIDKTIIGDFQKEQKKFVEEQHTKKSEA 20 AVPPWVDTNDEETIQQQILALSADKRNFLRDPPAGVQFNFDFDQMYPVALVML (SEQ ID NO:245); MRFALVPKLVKEEVFWRNYFYRVSLIKQSAQLTALAAQQQA AGKGGEEQ (SEQ ID NO:246); STSPGVSEFVSDAFDACNLNQEDLRKEMEQL VLDKKQEETAVLEEDSADWEKELQQELQEYEVVTESEKRDENWDK (SEO ID NO:247); SPWESRRVESKATSARCGLWGSGPRRRPASGMFRGLSSWLGLQQ 25 PVAGGGQPNGDAPPEQPS (SEQ ID NO:248); PVAGGGQPNGDAPPEQPSETV ESAEELQQAGDQELLHQAKDFGNYLFNFASAATKKITESVAE (SEO ID NO: 249); and/or FQKEQKKFVEEQHTKKSEAAVPPWVDTNDEETIQQQILALSADKR NFLRDPPAGVQFNFDFDQMYPVALVML (SEQ ID NO:250). Also preferred are 30 polynucleotide fragments encoding these polypeptide fragments.

This gene is expressed primarily in kidney pyramids and to a lesser extent in lung and other tissues of various types. This gene fluxes calcium in human aortic smooth muscle cells, and therefore is involved in signal transduction.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are

not limited to, renal and nervous disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the kidney and/or nervous system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., kidney, lung, brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution in kidney and lung and homology with sap47 indicates that the protein product has regulatory or direct functions in molecular exchange with body fluids and nervous system signaling. Polynucleotides and polypeptides corresponding to this gene are useful for treatment of disorders in kidney and nervous system.

FEATURES OF PROTEIN ENCODED BY GENE NO: 5

The translation product of this gene shares sequence homology with the mouse Ly-9.2 antigen which is thought to be an important cell surface marker in lymphoids, myeloids and hematopoietic progenitors. (See Accession No. gil198932.) Preferred polypeptide fragments comprise the amino acid sequence: PFICVARNPVSRNFSSPI LARKLCEGAA (SEQ ID NO:251); and/or KEDPANTVYSTVEIPKKMENPHSLLT MPDTPRL (SEQ ID NO:252). Also preferred are polynucleotide fragments encoding these polypeptide fragments. Based on homology, it is likely that this gene is also a cell surface marker, involved in hematopoiesis.

This gene is expressed primarily in activated macrophages, monocytes and T-cells and to a lesser extent in spleen and bone marrow.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune and hematopoietic disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and hematopoietic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., blood cells, and bone marrow, and cancerous and wounded

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tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:129 as residues: Lys-26 to Tyr-33. Arg-44 to Ile-49, Ser-53 to Lys-71, Lys-86 to Pro-91.

The tissue distribution and homology to Ly-9.2 surface immunoglobulin family indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis of immune and hematopoietic disorders. Polypeptides and polynucleotides corresponding to this gene are also be used as a marker for leukemia or a modulator of the functions of the cells of macrophage/monocyte or T-cell types.

FEATURES OF PROTEIN ENCODED BY GENE NO: 6

The translation product of this gene shares sequence homology with the *Drosophila* glutactin gene which is thought to be important in cell-cell interaction or cell-extracellular matrix contact.

This gene is expressed primarily in colon tissue, aorta endothelial cells and to a lesser extent in skin, breast tissue and T-cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of these tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, diseases of the gastrointestinal tract, vascular system or T-cell development. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the digestive system, cardiovascular system, and immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., colon, cardiovascular tissue, skin, mammary tissue, and blood cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to glutactin indicates that polynucleotides and polypeptides corresponding to this gene are useful for the development and maintenance of the integrity of the basal membrane in the gastrointestinal tract and

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cardiovascular system. The expression in T-cells also indicate the protein may be involved in T-cell adhesion, cell-cell interaction and development.

FEATURES OF PROTEIN ENCODED BY GENE NO: 7

The translation product of this gene shares sequence homology with MURF4 protein, an ATPase homolog, which is thought to be important in ATP hydrolysis.

This gene is expressed primarily in breast tissue.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, breast cancer and non-neoplastic breast diseases. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the breast tissue, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., mammary tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to MURF4 gene indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of neoplastic or non-neoplastic breast diseases because ATPase like protein may be involved in changed metabolic states of the breast.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 8

This gene shares homology to the alcohol dehydrogenase gene. Preferred polypeptide fragments comprise comprise the amino acid sequence: ASAVLLDLPNSG GEAQAKKLGNNCVFAPADVTSEKDVQTALALAKGKFGRVDVAVNCAGIAVAS KTYNLKKGQTHTLEDFQRVLDVNLMGTFNVIRLVAGEMGQNEPDQGGQRGVI INTASVAAFEGQVGQAAYSASKGGIVGMTLPIARDLAPIGIRVMTIAPGLFGTPL LTSLPEKVCNFLASQVPFPSRLGDPAEYAHLVQAIIENPFLNGEVIRLDGAIRMQ P (SEQ ID NO:253); and/or SVAAFEGQVGQAAYSASKGGIVGMTLPIA (SEQ ID NO:254). Polynucleotides encoding these fragements are also encompassed by the invention. Other groups have also recently cloned this gene, recognizing its homology to alcohol dehydrogenase. (See Accession No. 1778355.) Moreover, a second group

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recently cloned the mouse homologue of this gene. (See Accession No. 2078284.) They found that the mouse homologue binds to amyloid beta-peptide and mediates neurotoxicity in Alzheimer's disease, calling the protein ERAB. This gene maps to chromosome X, and therefore can be used in linkage analysis as a marker for chromosome X. Therefore, mutations in the translated product of this gene may be involved in Alzheimer's disease in humans, as well as other sex linked diseases. This gene can be used as a diagnostic marker for these diseases.

Preferred epitopes include those comprising a sequence shown in SEQ ID NO:132 as residues: Arg-45 to Ser-53.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 9

The translation product of this gene shares week sequence homology with rat N-methyl-D-aspartate receptor subunit and other proline-rich proteins which are thought to be important in neurotransmission or protein-protein intereaction.

This gene is expressed primarily in synovial hypoxia and to a lesser extent in ovary, senescent cells and brain.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, synovial hypoxia. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the synovia and brain, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., synovial tissue, ovary and other reproductive tissue, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution in synovial hypoxia and nerve tissues, and homology to N-methyl-D-aspartate receptor subunit and other proline-rich proteins indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and intervention of synovial hypoxia and other synovial disorders.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 10

This gene is expressed primarily in prostate and to a lesser extent in placenta and ovary.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, male and female infertility, cancer, and other hyperproliferative disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the reproductive system and neoplasia, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., prostate, placenta, ovary and other reproductive tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:134 as residues: Pro-17 to Met-23, Ala-30 to Trp-38, Ile-49 to Trp-54, Lys-68 to Gly-74, Thr-93 to Gly-99, Met-126 to Glu-132, Gly-173 to Ser-178, Lys-205 to Tyr-214.

The tissue distribution of this gene in the prostate, placenta and ovary indicates that this gene product is useful for treatment/diagnosis of male or female infertility, endocrine disorders, fetal deficiencies, ovarian failure, amenorrhea, ovarian cancer, benign prostate hyperplasia, prostate cancer, and other forms of cancer of the reproductive system.

FEATURES OF PROTEIN ENCODED BY GENE NO: 11

This gene is expressed primarily in the thyroid and to a lesser extent in the pineal gland. This gene maps to chromosome 10, and therefore can be used as a marker in linkage analysis for chromosome 10. Preferred polypeptide fragments comprise the amino acid sequence: HPIEWAINAATLSQFY (SEQ ID NO:256); CWIKYCLTLMQN AQLSMQDNIG (SEQ ID NO:257); KVSYLRPLDFEEARELFLLGQHYVF (SEQ ID NO:258); MERRCKMHKRXIAMLEPLTVDLNPQ (SEQ ID NO:259); and/or SHIV KKINNLNKSALKY YQLFLD (SEQ ID NO:260). Also preferred are polynucleotides encoding these polypeptide fragments.

Therefore, polynucleotides and polypeptides of the invention are useful as

reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune, thyroid and pineal gland disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and endocrine systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., thyroid and pineal gland, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:135 as residues: Ser-2 to Ser-8, Thr-38 to Arg-44.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treating/detecting immune disorders such as arthritis, asthma, immune deficiency diseases (e.g., AIDS), and leukemia, as well as treating/detecting thymus disorders (e.g., Graves Disease, lymphocytic thyroiditis, hyperthyroidism, and hypothyroidism), and treating/detecting pineal gland disorders (e.g., circadian rhythm disturbances associated with shift work, jet lag, blindness, insomnia and old age).

FEATURES OF PROTEIN ENCODED BY GENE NO: 12

This gene is expressed primarily in lung and tonsils.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, pulmonary or immune disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the pulmonary and immune systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., pulmonary tissue, and tonsils, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily

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fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:136 as residues: Glu-28 to Gly-49.

The tissue distribution of this gene only in lung indicates that it could play a role in the treatment/detection of lung lymphoma or sarcoma formation, pulmonary edema and embolism, bronchitis and cystic fibrosis. Its expression in tonsils indicates a potential role in the treatment/detection of immune disorders such as arthritis, asthma, immune deficiency diseases (e.g., AIDS), and leukemia, in addition to the treatment/detection of tonsillitis.

FEATURES OF PROTEIN ENCODED BY GENE NO: 13

This gene is expressed primarily in lymphoid, myeloid and erythroid cells. Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, hematopoietic and immune disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the hematopoietic and immune systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., blood cells, myeloid cells, and bone marrow, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level

The predominant tissue distribution of this gene in hematopoietic cell types indicates that the gene could be important for the treatment or detection of immune or hematopoietic disorders including arthritis, asthma, immunodeficiency diseases and leukemia. Preferred embodiments of the present invention are polypeptide fragments comprising the amino acid sequence: FTHLSTCLLSLLLVRMSGFLLLARASPSI CALDSSCFVEYCSSYSSSCFLHQHFPSLLDHLCQ (SEQ ID NO:261); or FLLL ARASPSICALDSSCFVQEY (SEQ ID NO:262). Also preferred are polynucleotide fragments encoding these polypeptide fragments.

in healthy tissue or bodily fluid from an individual not having the disorder.

FEATURES OF PROTEIN ENCODED BY GENE NO: 14

This gene is homologous to the *Drosophila Regena* (*Rga*) gene. (See Accession No. 1658504.) This *Drosophila* gene is thought to be a homolog of the global negative

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transcriptional regulator NOT2 (CDC36) from yeast, which modifies gene expression and suppresses position effect variegation. Preferred polypeptide fragments comprise the amino acid sequence: PDGRVTNIPQGMVTDQFGMIGLLTFIRAAETDPGMVHL ALGSDLTTLGLNLNS (SEQ ID NO:263); VHLALGSDLTTLGLNLNSPENLYP (SEQ ID NO:265); EDLLFYLYYMNGGDVLQLLAAVELFNRDWRYHKEERVWI TR (SEQ ID NO:264); and/or HNEDFPALPGS (SEQ ID NO:266).

This gene is expressed primarily in placenta and to a lesser extent in infant brain.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, neurodegenerative and developmental disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the neurological system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., placenta, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:138 as residues: Leu-9 to Tyr-15, Asp-34 to Gln-46, Pro-51 to Asp-57, Gly-88 to Thr-104, Thr-123 to Ser-128.

The tissue distribution of this gene indicates that it could be used in the detection and/or treatment of neurological disorders such as such as Alzheimer's Disease, Parkinson's Disease, Huntington's Disease, schizophrenia, mania, dementia, paranoia, obsessive compulsive disorder, and panic disorder.

30 FEATURES OF PROTEIN ENCODED BY GENE NO: 15

This gene is expressed primarily in adrenal gland tumor and osteoclastoma.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, endocrine and bone disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for

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differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the endocrine system and in bone, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., adrenal gland, and bone, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:139 as residues: Ile-52 to Trp-57.

The tissue distribution of this gene indicates that it may be involved in the treatment and/or detection of adrenal gland tumors, osteosarcomas, endocrine disorders and bone disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 16

The translation product of this gene shares sequence homology with the FK506 binding protein, a protein which plays an important role in immunosupression. (See Accession No. M75099.) Specifically, a 12-kDa FK506-binding protein (FKBP-12) is a cytosolic receptor for the immunosuppressants FK506 and rapamycin. (See, Proc. Natl. Acad. Sci. 88: 6677-6681 (1991).) Thus, based on homology, it is likely that this gene also has immunosuppression activity. Preferred polypeptides comprise the amino acid sequence: GRIIDTSLTRDPLVIELGQKQVIPGLEQSLLDMCVGEKRRAIIPSH LAYGKRGFPPSVPADAVVQYDVELIALIR (SEQ ID NO:267); and/or IHYTGSLV DGR IIDTS (SEQ ID NO:268). Also preferred are the polynucleotide fragments encoding these polypeptides.

This gene is expressed primarily in melanocytes.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cancer and other hyperproliferative disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system and cancer, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., melanocytes, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to

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the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:140 as residues: Ala-118 to Phe-124, Arg-178 to Lys-201.

The tissue distribution and homology to the FK506 binding proteins which are believed to a role in immunosupression mediated by the immunosupressant drugs rapamycin and cyclosporin, indicates that this gene could serve as a novel target for the identification of novel immunosupressant drugs.

10 FEATURES OF PROTEIN ENCODED BY GENE NO: 17

The translation product of this gene shares sequence homology with the rat calcium-activated potassium channel rSK3, which is thought to be important in regulating vascular tone. (See Accession No. gil2564072, gil1575663, and gil1575661.) Although homologous to these proteins, this gene contains an 18 amino acid insert, not previously identified in the homologs. Preferred polypeptide fragments comprise the amino acid sequence: CESPESPAQPSGSSLPAWYH (SEQ ID NO:269). Also preferred are the polynucleotide fragments encoding these polypeptides.

This gene is expressed primarily in B-cells, frontal cortex and endothelial cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cardiovascular (hyper/hypotension, asthma, pulmonary edema, pneumonia, heart disease, restenosis, atherosclerosis, stoke, angina and thrombosis) and neurological disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the cardiovascular and nervous systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., blood cells, brain and other tissue of the nervous system, and endothelium, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:141 as residues: Glu-72 to Gly-82, His-90 to Val-95, Gln-168 to Lys-174, Val-202 to Ser-212.

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The tissue distribution and homology to calcium-activated potassium channels indicates that polynucleotides and polypeptides corresponding to this gene are useful for the diagnosis and treatment of vascular disorders (hyper/hypotension, athesma, pulmonary edema, pneumonia, heart disease, restenosis, atherosclerosis, stoke, angina and thrombosis).

FEATURES OF PROTEIN ENCODED BY GENE NO: 18

This gene is expressed primarily in smooth muscle and to a lesser extent in brain (amygdala, corpus colosum, hippocampus).

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cardiovascular (hypertension, heart disease, athesma, pulmonary edema, restenosis, atherosclerosis, stoke, angina, thrombosis, and wound healing), and neurological disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the cardiovascular and neurological systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., smooth muscle, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:142 as residues: Lys-43 to Arg-49, Tyr-58 to Glu-65.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of cadiovascular disorders (hypertension, heart disease, athesma, pulmonary edema, restenosis, atherosclerosis, stoke, angina, thrombosis, and wound healing). Expression in brain indicates a role in the treatment and diagnosis of behavioral or neurological disorders, such as depression, schizophrenia, Alzheimer's disease, mania, dementia, paranoia, and addictive behavior.

35 FEATURES OF PROTEIN ENCODED BY GENE NO: 19

This gene is expressed primarily in T-cells (Jurkats, resting, activated, and

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anergic T-cells), endothelial cells, pineal gland, and to a lesser extent in a variety of other tissues and cell types. Preferred polypeptide fragments comprise the amino acid sequence: EEAGAGRRCSHGGARPAGLGNEGLGLGGDPDHTDTGSRSKQRINN WKESKHKVIMASASARGNQDKDAHFPPPSKQSLLFCPKSKLHIHRAEISK (SEQ ID NO:270); and/or SKQRINNWKESKHKVIMASASAR (SEQ ID NO:271). Also preferred are the polynucleotide fragments encoding these polypepides.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, inflammation, immune and cardiovascular disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune. neurological and vascular systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, endothelial cells, and pineal gland, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:143 as residues: Phe-71 to Arg-76, Pro-82 to His-87, Glu-103 to Ala-111.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the diagnosis and treatment of immune disorders including: leukemias, lymphomas, auto-immune, immuno-supressive (e.g. transplantation) and immunodeficiencies (e.g. AIDS) and hematopoietic disorders. In addition, expression in the pineal gland might suggest a role in the diagnosis of specific brain tumors and treatment of neurological disorders. Endothelial cell expression might suggest a role in cadiovascular or respiratory/pulmonary disorders or infections (athesma, pulmonary edema, pneumonia).

FEATURES OF PROTEIN ENCODED BY GENE NO: 20

This gene is expressed primarily in brain and embryo and to a lesser extent in leukocytes. This gene maps to chromosome 15, and therefore can be used as a marker in linkage analysis to chromosome 15.

Therefore, polynucleotides and polypeptides of the invention are useful as

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reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, developmental and neurological disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the central nervous and immune systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:144 as residues: Met-1 to Gly-8.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of immune disorders including: leukemias, lymphomas, auto-immune, immuno-supressive (e.g. transplantation) and immunodeficiencies (e.g. AIDS) and hematopoietic disorders. The expression in the brain -- and in particular the fetal brain -- would suggest a possible role in the treatment and diagnosis of developmental and neurodegenerative diseases of the brain and nervous system (depression, schizophrenia, Alzheimer's disease, mania, dementia, paranoia, and addictive behavior).

FEATURES OF PROTEIN ENCODED BY GENE NO: 21

This gene is expressed primarily in brain, kidney, lung, liver, spleen, and a variety of leukocytes (especially T-cells) and to a lesser extent in a variety of other tissues and cell types.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, leukemias, lymphomas, autoimmune, immunosuppressive, and immunodeficiencies, hematopoietic disorders, as well as renal disorders, and neoplasms. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the renal, pulmonary, immune, and central nervous systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g.,

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brain and other tissue of the nervous system, kidney, pulmonary tissue, liver, spleen, and blood cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of renal conditions, such as acture renal failure, kidney fibrosis, and kidney tubule regeneration. The expression in leukocytes and other immune tissues indicates a role in immune disorders including: leukemias, lymphomas, auto-immune, immuno-supressive (e.g. transplantation) and immunodeficiencies (e.g. AIDS) and hematopoietic disorders. The expression in the brain -- and in particular the fetal brain -- indicates a possible role in the treatment and diagnosis of developmental and neurodegenerative diseases of the brain and nervous system (depression, schizophrenia, Alzheimer's disease, mania, dementia, paranoia, and addictive behavior).

FEATURES OF PROTEIN ENCODED BY GENE NO: 22

This gene is expressed primarily in skin (fetal epithelium, keratinocytes and skin). This gene also maps to chromosome 19, and therefore can be used in linkage analysis as a marker for chromosome 19.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, skin cancers (e.g., melanomas), eczema, psoriasis or other disorders of the skin. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the skin, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., skin and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:146 as residues: Pro-28 to Glu-35, Ser-39 to Phe-44, Ala-94 to Gln-99.

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The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of skin cancers (e.g., melanomas), eczema, psoriasis or other disorders of the skin.

5 FEATURES OF PROTEIN ENCODED BY GENE NO: 23

This gene maps to chromosome 11. Another group recently isolated this same gene, associating the sequence to the region thought to harbor the gene involved in Multiple Endocrine Neoplasia Type 1, or MEN 1. (See Accession No. 2529721 and Genome Res. 7(7), 725-735 (1997), incorporated herein by reference in its entirety.) Preferred polypeptide fragments comprise the amino acid sequence: LFHWACLNERA AOLPRNTAXAGYQCPSCNGPS (SEQ ID NO:272).

This gene is expressed primarily in epididymus, pineal gland, T-cells, as well as fetal epithelium, lung and kidney.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune, metabolic mediated disorders, and MEN. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, renal, neurological and pulmonary systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., epididymus and other reproductive tissue, pineal gland, T-cells and other blood cells, epithelium, lung, and kidney, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of developmental deficiencies or abnormalities as well as a host of different disorders which arise as a result of conditions in the indicated tissues or cell types. An area of particular interest is in the treatment and diagnosis of immune disorders including: leukemias, lymphomas, auto-immune, immuno-supressive (e.g. transplantation) and immunodeficiencies (e.g. AIDS) and hematopoietic disorders. The expression in the brain, and in particular the fetal brain, would suggest a possible role in the treatment and diagnosis of

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developmental and neurodegenerative diseases of the brain and nervous system (depression, schizophrenia, Alzheimer's disease, mania, dementia, paranoia, and addictive behavior). Respiratory/pulmonary disorders, such as athesma, pulmonary edema are also potential therapeutic areas, as well as renal conditions such as acute renal failure, kidney fibrosis and kidney tubule regeneration. Moreover, this gene can be used in the treatment and/or detection of MEN I.

FEATURES OF PROTEIN ENCODED BY GENE NO: 24

This gene is expressed primarily in fetal spleen.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, leukemia, lymphoma, AIDS, hematopoeitic disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and hematopoietic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., spleen and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of immune disorders including: leukemias, lymphomas, auto-immune, immuno-supressive (e.g. transplantation) and immunodeficiencies (e.g. AIDS) and hematopoietic disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 25

A closely related homolog of this gene was recently cloned by another group,

calling the gene CDO, an oncogene-, serum-, and anchorage-regulated member of the
Ig/fibronectin type III repeat family. (See Accession No. 2406628, and J. Cell Biol.
138(1): 203-213 (1997), herein incorporated by reference in its entirety.) Preferred
polypeptide fragments comprise the amino acid sequence: FYIYYRPTDSDNDSDYKK
DMVEGDKYWHSISHLQPETSYDIKMQCFNEGGESEFSNVMICETKARKSSGQP
GRLPPPTLAPPQPPLPETIERPVGTGAMVARSSDLPYLIVGVVLGSIVLIIVTFIPF
CLWRAWSKQKHTTDLGFPRSALPPSCPYTMVPLGGLPGHQAVDSPTSVASVD

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GPVLM (SEQ ID NO:273); or YIYYRPTDSDNDSDYKKDMVEGDKYWHSISHLQ PETSYDIKMQCFNEGGESEFSNVMICETKARKS (SEQ ID NO:274).

This gene is expressed primarily in fetal lung and kidney, human embryo and osteoclastoma stromal cells and to a lesser extent in a variety of other tissues and cell types.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, developmental disorders and cancers, as well as pulmonary and renal disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the respiratory/pulmonary, skeletal and renal systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., lung, kidney, embryonic tissue, and bone cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:149 as residues: Thr-5 to Pro-18, Ala-76 to Thr-84.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the detection and treatment of: osteoperosis, fracture, osteosarcoma, ossification, and osteonecrosis, as well as respiratory/pulmonary disorders, such as athesma, pulmonary edema, and renal conditions such as acute renal failure, kidney fibrosis and kidney tubule regeneration.

FEATURES OF PROTEIN ENCODED BY GENE NO: 26

This gene is homologous to the HIV envelope glycoprotein. (See Accession No. 2641463.) Preferred polypeptide fragments comprise the amino acid sequence: NVRALLHRMPEPPKINTAKFNNNKRKNLSL (SEQ ID NO:275).

This gene is expressed primarily in pineal gland and skin, and to a lesser extent in lung.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are

not limited to, neurological and behavior disorders: respiratory/pulmonary disorders, such as athesma, pulmonary edema; skin conditions such as eczema, psoriasis, acne and skin cancer, as well as AIDS. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the central nervous and respiratory systems, as well as skin and AIDS, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., blood cells, pineal gland, epidermis, and pulmonary tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:150 as residues: Gln-15 to Gln-20.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of conditions which affect the above tissues, such as: skin cancer, eczema, psoriasis, acne, athesma, pulmonary edema, neuro-degenerative or developmental disorders such as Alzheimer's, depression, schizophrenia, dementia, and AIDS.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 27

Preferred polypeptide encoded by this gene comprise the following amino acid sequence: NTNQREALQYAKNFQPFALNHQKDIQVLMGSLVYLRQGIENSPYVHL LDANQWADICDIFTRDACALLGLSVESPLSVSFSAGCVALPALINIKAVIEQRQC TGVWNQKDELPIEVDLGKKCWYHSIFACPILRQQTTDNNPPMKLVCGHIISRD ALNKMFNGSKLKCPYCPMEQSPGDAKQIFF (SEQ ID NO:276). Polynucleotides encoding such polypeptides are also provided as are complementary polynucleotides thereto.

This gene is expressed primarily in liver (adult and fetal) and spleen tissue, and to a lesser extent in placenta, T helper cells, kidney tumor, ovarian tumor, melanocytes and fetal heart.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune and developmental diseases and disorders and liver diseases such as liver cancer. Similarly, polypeptides and antibodies directed to these

polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, circulatory and hematopoietic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., liver, spleen, placenta, blood cells, kidney, ovary and other reproductive tissue, melanocytes, and heart, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that the protein products of this gene are useful for study, diagnosis and treatment of growth, hematopoietic and immune system disorders particularly related to the liver.

FEATURES OF PROTEIN ENCODED BY GENE NO: 28

The translation product of this gene shares sequence homology with prostaglandin transporter which is thought to be important in metabolic and endocrine disorders. See, for example, Gastroenterology Oct:109(4):1274-1282 (1995). Preferred polypeptides encoded by this gene comprise the following amino acid sequence: SYLSACFAGCNSTNLTGCACLTTVPAENATVVPGKCPSPGCQEAFLTFLCVMCI CSLIGAMARHP (SEQ ID NO:277); and/or PSVIILIRTVSPELKSYALGVLFLLLRL LGFIPPPLIFGAGIDSTCLFWSTFCGEQGACVLYDNVVYRYLYVSIAIALKSFAFI (SEQ ID NO:278).

This gene is expressed primarily in hematopoietic and brain tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, metabolic, immune and endocrine diseases and disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the metabolic, immune and endocrine systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., endocrine tissue, hematopoietic tissue, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to

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the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to prostaglandin (and anion) transporter indicates that polynucleotides and polypeptides corresponding to this gene are useful for study, diagnosis and treatment of endocrine, metabolic, immune and kidney disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 29

This gene is expressed primarily in early stage human lung.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, growth and respiratory disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the developmental and respiratory systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., pulmonary tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:153 as residues: Val-50 to Trp-55.

The tissue distribution indicates that the protein products of this gene are useful for study, diagnosis and treatment of respiratory and growth diseases and disorders.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 30

The translation product of this gene shares sequence homology with human DNA helicase which is thought to be important in accurate and complete DNA replication in creation of new cells. Preferred polypeptides encoded by this gene comprise the following amino acid sequence: QSLFTRFVRVGVPTVDLDAQGRARA SLCXXYNWRYKNLGNLPHVQLLPEFSTANAGLLYDFQLINVEDFQGVGESEPN PYFYQNLGEAEYVVALFMYMCLLGYPADKISILTTYNGQKHLIRDIINRRCGNN PLIGRPNKVTTVDRFQGQQNDYILLSLVRTRAVGHLRDVRRLVVAMSRAR (SEQ ID NO:279); and/or LVKEAKIIAMTCTHAALKRHDLVKLGFKYDNILMEE AAQILEIETFIPLLLQNPQDGFSRLKRWIMIGDHHQLPPVI (SEQ ID NO:280).

This gene is expressed primarily in testes tumor and to a lesser extent in adrenal

gland tumor and placenta.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cancers and endocrine/growth disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the endocrine, developmental, and reproductive systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., testes and other reproductive tissue, adrenal gland, and placenta, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to DNA helicase indicates that the protein products of this gene are useful for study, treatment, and diagnosis of many cancer types, including testicular cancer; as well as disorders involving endocrine function and normal growth and development.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 31

The translation product of this gene shares sequence homology with BID-apoptotic death gene (mouse), Genbank accession no. PID g1669514, which is thought to be important in programmed cell death.

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This gene is expressed primarily in jurkat membrane bound polysomes and activated neutrophils and to a lesser extent in endothelial cells and human cerebellum.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cancers and other proliferative disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., blood cells, endothelium, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma,

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urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:155 as residues: Glu-4 to Leu-11, Cys-28 to Arg-35, Gln-50 to His-66, Glu-73 to Gln-79, Gly-94 to Ser-100, Arg-114 to Asp-126, Pro-139 to Lys-146.

The tissue distribution and homology to BID-apoptotic death gene indicates that the protein products of this gene are useful for study of cell death, and treatment and diagnosis of proliferative disorders and cancers. Apoptosis - programmed cell death - is a physiological mechanism involved in the deletion of peripheral T lymphocytes of the immune system, and its dysregulation can lead to a number of different pathogenic processes. Diseases associated with increased cell survival, or the inhibition of apoptosis, include cancers (such as follicular lymphomas, carcinomas with p53 mutations, and hormone-dependent tumors, such as breast cancer, prostrate cancer, Kaposiís sarcoma and ovarian cancer); autoimmune disorders (such as systemic lupus erythematosus and immune-related glomerulonephritis rheumatoid arthritis) and viral infections (such as herpes viruses, pox viruses and adenoviruses), inflammation; graft vs. host disease, acute graft rejection, and chronic graft rejection. Diseases associated with increased apoptosis include AIDS; neurodegenerative disorders (such as Alzheimer's disease, Parkinson's disease, Amyotrophic lateral sclerosis, Retinitis pigmentosa, Cerebellar degeneration); myelodysplastic syndromes (such as aplastic anemia), ischemic injury (such as that caused by myocardial infarction, stroke and reperfusion injury), toxin-induced liver disease (such as that caused by alcohol), septic shock, cachexia and anorexia. Thus, the invention provides a method of enhancing apoptosis in an individual by treating the individual with a polypeptide encoded by this gene.

FEATURES OF PROTEIN ENCODED BY GENE NO: 32

The translation product of this gene shares sequence homology with human fructose transporter which is thought to be important in normal metabolic function and activity.

This gene is expressed primarily in T-cell lymphoma.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are

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not limited to, leukemia and other cancers, and metabolic disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the hematopoietic, lymph and metabolic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:156 as residues: Pro-22 to Gly-48, Ser-54 to Pro-61.

The tissue distribution indicates that the protein products of this gene are useful for study of mechanisms leading to cancer, treatment and diagnosis of cancerous and pre-cancerous conditions; as well as the study and treatment of various metabolic diseases and disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 33

This gene is expressed primarily in human meningima.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, inflammation and other disorders of the CNS. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the CNS and immune systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., meningima and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:157 as residues: Asn-23 to Pro-31.

The tissue distribution indicates that the protein products of this gene are useful for study, diagnosis and treatment of disorders of the CNS and inflammatory responses.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 34

This gene is expressed primarily in activated monocytes and wound healing tissues and to a lesser extent in fetal epithelium.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune and inflammatory disorders and wound healing and tissue repair dysfunctions. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, epithelial and gastrointestinal systems, and healing wounds, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., monocytes and other blood cells, and epithelium, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:158 as residues: Ala-28 to Ala-33, Gly-35 to Glu-45.

The tissue distribution indicates that the protein products of this gene are useful for diagnosis, study and treatment of immune and inflammatory disorders and wound healing dysfunctions.

25 FEATURES OF PROTEIN ENCODED BY GENE NO: 35

This gene is expressed primarily in human osteosarcoma and prostate cancer.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, skeletal and neoplastic conditions such as bone and prostate cancer. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and skeletal systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., bone, and prostate, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial

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fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:159 as residues: Ser-14 to Gly-22, Leu-37 to Gln-43.

The tissue distribution indicates that the protein products of this gene are useful for diagnosis and treatment of skeletal disorders and cancer.

FEATURES OF PROTEIN ENCODED BY GENE NO: 36

This gene encodes a protein which is highly homologous to a protein called congenital heart disease protein 5, presumably implicated in congenital heart disease (see Genbank PID g2810996).

This gene is expressed primarily in Hodgkin's lymphoma, erythroleukemia cells, and TNF activated synovial fibroblasts, to a lesser extent in ovarian cancer, cerebellum, spleen, fetal liver and placenta and finally to a lesser extent in various other mesenchymal tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cancer, immune, hematopoietic and cardiovascular disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, hematopoietic and cardiovascular systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., heart and other cardiovascular tissue, lymphoid tissue, blood cells, bone marrow, ovary and other reproductive tissue, brain and other tissue of the nervous system, spleen, liver, and mesenchymal tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:160 as residues: Lys-41 to Met-49, Gln-54 to Glu-59, Glu-76 to Thr-88.

The homology of this gene and translation product to congenital heart disease protein 5 indicates a role for this protein in the diagnosis, prognosis and/or treatment of

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heart disease or other cardiovascular related disorders. In addition, predominant expression in cells associated with the immune and hematopoetic system indicates a role for this protein in the treatment, diagnosis and/or prognosis of immune and autoimmune diseases, such as lupus, transplant rejection, allergic reactions, arthritis, asthma, immunodeficiency diseases, leukemia, AIDS, thymus disorders such as Graves Disease, lymphocytic thyroiditis, hyperthyroidism and hypothyroidism, graft versus host reaction, graft versus host disease, transplant rejection, myelogenous leukemia, bone marrow fibrosis, and myeloproliferative disease. The protein could also be used to enhance or protect proliferation, differentiation and functional activation of hematopoietic progenitor cells such as bone marrow cells, which could be useful for cancer patients undergoing chemotherapy or patients undergoing bone marrow transplantation. The protein may also be useful to increase the proliferation of peripheral blood leukocytes, which could be useful in the combat of a range of hematopoietic disorders including immunodeficiency diseases, leukemia, and septicemia.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 37

This gene is expressed primarily in ovarian cancer.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, urogenital neoplasias. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the reproductive system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., ovary and other reproductive tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:161 as residues: Asn-22 to Asn-27.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for study, diagnosis and treatment of ovarian and other tumors.

FEATURES OF PROTEIN ENCODED BY GENE NO: 38

The translation product of this gene shares sequence homology with zinc finger proteins.

This gene is expressed primarily in various fetal, cancer, and endothelial lines.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune and growth disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the cardiovascular system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., fetal tissue, and endothelial cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that the protein products of this gene are useful for study, diagnosis and treatment of immune and developmental conditions and cancer.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 39

This gene is expressed primarily in fetal, infant, and adult brain and to a lesser extent in other brain and endocrine organs and blastomas.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, brain tumors and neurodegenerative conditions. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the nervous and endocrine systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., brain and other tissue of the nervous system, endocrine tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the

disorder.

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The tissue distribution indicates that the protein products of this gene are useful for the study, diagnosis and treatment of brain cancer and other neurological disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 40

The translation product of this gene shares sequence homology with vesicular glycoproteins and lectins. Preferred polypeptides encoded by this gene comprise the following amino acid sequence: DTYPNEEKQQERVFPXXSAMVNNGSLSYDHER DGRPTELGGCXAIVRNLHYDTFLVIRYVKRHLTIMMDIDGKHEWRDCIEVPGV RLPRGYYFGTSSITGDLSDNHDVISLKLFELTVERTPEEE (SEQ ID NO:281); and/or LKREHSLSKPYQGVGTGSSSLWNLMGNAMVMTQYIRLTPDMQSKQGA LWNRVPCFLRDWELQVHFKIHGQGKKNLHGDGLAIWYT (SEQ ID NO:282).

This gene is expressed primarily in infant brain and to a lesser extent in various normal and transformed neural, endocrine, and immune organs.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, neurological and neurodevelopmental conditions. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the nervous and hormonal systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., brain and other tissue of the nervous system, endocrine tissue, and tissue and cells of the immune system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:164 as residues: Pro-64 to Gly-71, Gly-94 to Leu-100, Thr-110 to Pro-116, Thr-135 to Arg-145, Glu-164 to Glu-171, Asp-204 to Asp-211, Arg-253 to His-261, Asn-312 to Tyr-323.

The tissue distribution indicates that the protein products of this gene are useful for the study, diagnosis and treatment of mental retardation and other neurological disorders and neoplasias.

FEATURES OF PROTEIN ENCODED BY GENE NO: 41

This gene displays homology to the glycosyltransferase family, which catalyze the addition of sialic acids to carbohydrate groups which are present on glycoproteins.

This gene is expressed primarily in smooth muscle and to a lesser extent in pineal gland, fetal liver, and infant brain.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, gastrointestinal injury, inflammatory and neurodegenerative conditions. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and nervous systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., smooth muscle, pineal gland, liver, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:165 as residues: Ser-12 to Trp-21, Arg-24 to Pro-32, Asp-73 to Lys-82, Lys-90 to Ala-97.

The tissue distribution indicates that the protein products of this gene are useful for the study, diagnosis and treatment of neurodegenerative and growth disorders and gastrointestinal repair.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 42

The translation product of this gene shares sequence similarity with metallothionein polypeptides. See, for example, Proc. Natl. Acad. Sci. U S A 1992 Jul 15:89(14):6333-6337. Metallothioneins are believed to inhibit neuronal survival among other biological functions. Based on the sequence similarity (especially the conserved cysteine motifs characteristic of the metallothionein family) the translation product of this gene is expected to share certain biological activities with other members of the metallothionein polypeptide family. Preferred polypeptides encoded by this gene comprise the following amino acid sequence: PGTLQCSALHHDPGCANCSRFCRD CSPPACQC (SEQ ID NO:283).

This gene is expressed exclusively in placenta and fetal liver.

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Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, hematopoietic and immune disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the reproductive and immune systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., placenta, liver, brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to metallothionien indicates that the protein products of this gene are useful for diagnosis and treatment of immune and hematopoietic system disorders and neurological diseases, especially in fetal development.

20 FEATURES OF PROTEIN ENCODED BY GENE NO: 43

Preferred polypeptides encoded by this gene comprise the following amino acid sequence: FLYDVLMXHEAVMRTHQIQLPDPEFPS (SEQ ID NO:284).

This gene is expressed primarily in T-cells and synovial tissue.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune system disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., synovial tissue, and T-cells and other blood cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

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The tissue distribution indicates that the protein products of this gene are useful for treatment and diagnosis of disorders of the immune system.

FEATURES OF PROTEIN ENCODED BY GENE NO: 44

The translation product of this gene shares sequence similarity with several methyltransferases (e.g., see Genbank gil1065505).

This gene is expressed primarily in ovary, thymus, infant adrenal gland, tissues of the nervous system and the hematopoietic tissue, and to a lesser extent in adipose tissue and many other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, disorders of the reproductive system, the endocrine system, the hematopoietic system and the CNS. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, endocrine, CNS and reproductive system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., ovary and other reproductive tissue, thymus, adrenal gland, brain and other tissue of the nervous system, hematopoietic tissue, and adipose tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:168 as residues: Ser-3 to Gly-12, Asp-19 to Arg-31, Tyr-70 to Tyr-77, Asn-130 to Lys-140, Pro-165 to Gln-170, Pro-192 to Lys-199, Leu-216 to Glu-227, Glu-254 to Phe-281.

The tissue distribution and homology to methyltransferase indicates that the protein products of this gene are useful for diagnosis and treatment of disorders of the CNS, the hematopoietic system and reproductive organs and tissues. For example, the abundant expression in the ovary may indicate that the gene product can be used as a hormone with either systemic or reproductive functions; as growth factors for germ cell maintenance and in vitro culture; as a fertility control agent; remedy for sexual dysfunction or sex development disorders; diagnostics/treatment for ovarian tumors, such as serous adenocarcinoma, dysgerminoma, embryonal carcinoma,

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choriocarcinoma, teratoma, etc; The expression in thymus may indicate its utilities in T-cell development and thus its applications in immune related medical conditions, such as infection, allergy, immune deficiency, tissue/organ transplantation, etc.

FEATURES OF PROTEIN ENCODED BY GENE NO: 45

The translation product of this gene shares sequence homology with cytochrome C oxidase which is thought to be important in metabolic function of cells. This gene has now recently been published as estrogen response gene. See Genbank accession no. AB007618 and Mol. Cell. Biol. 18 (1), 442-449 (1998). See also J Immunol. Mar 1:154(5): 2384-2392 (1995), where the mouse homologue was published and implicated in siliocis.

This gene is expressed primarily in adipose tissue, kidney and fetal brain and to a lesser extent in several other tissues and organs.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, metabolic diseases involving especially adipose tissue, brain and kidney. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the CNS and vascular system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., adipose tissue, kidney, brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:169 as residues: Thr-5 to Ser-14.

The tissue distribution and homology to cytochrome C oxidase, estrogen response gene product and siliocis related gene product indicates that the protein products of this gene are useful for diagnosis and treatment of metabolic disorders in the CNS, adipose tissue and kidney, particularly siliocis.

FEATURES OF PROTEIN ENCODED BY GENE NO: 46

The translation product of this gene shares sequence homology with reticulocalbin. See, for example, J. Biochem. 117 (5), 1113-1119 (1995). Based on the

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sequence similarity, the translation product of this gene is expected to share certain biological activities with reticulocalbin, e.g., Ca++ binding activities. This gene product is sometimes hereinafter referred to as "Reticulocalbin-2".

This gene is expressed primarily in breast, endothelial cells, synovial, heart and smooth muscle cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, diseases of the breast, vascular and skeletal/cardiac muscular system. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the breast, vascular and skeleto-muscular system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., mammary tissue, endothelial cells, synovial tissue, heart and other cardiovascular tissue, and smooth muscle, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:170 as residues: Gly-16 to Arg-32, Ala-42 to Asn-50, Glu-66 to Gln-76, Arg-85 to Gly-94, Thr-108 to Asp-115, Trp-121 to Gly-130. Leu-137 to His-144, Glu-155 to Lys-161, Asp-175 to Ser-180, Glu-209 to Gly-217, Glu-232 to Glu-237, Thr-243 to Asp-261, Glu-287 to Arg-295.

The tissue distribution indicates that the protein products of this gene are useful for diagnosis and treatment of diseases of the vascular and skeletal/cardiac muscular system. The homology of the gene with reticulocalbin indicates its biological function in regulating calcium store, a particularly important function in muscular cell types. The gene expression in the heart may indicate its utilities in diagnosis and remedy in heart failure, ischemic heart diseases, cardiomyopathy, hypertension, arrhythmia, etc. The abundant expression in the breast may indicate its applications in breast neoplasia and breast cancers, such as fibroadenoma, papillary carcinoma, ductal carcinoma, Pagetís disease, medullary carcinoma, mucinous carcinoma, tubular carcinoma, secretory carcinoma and apocrine carcinoma; juvenile hypertrophy and gynecomastia, mastitis and abscess, duct ectasia, fat necrosis and fibrocystic diseases, etc.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 47

The translation product of this gene shares weak sequence homology with H+-transporting ATP synthase which is thought to be important in cell metabolism or signal transduction.

This gene is expressed only in testis.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of some types of diseases and conditions. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the brain and hematopoietic tissues, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., testes and other reproductive tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

Since only one out of about a million expressed sequence tag is found in testes indicates that its expression is low and selectively in testes. Since some of the genes only expressed in testes are usually expressed in brain or in certain induced hematopoietic cells/tissues, it is speculated that this gene to be expressed in brain or hematopoietic cells/tissues and is useful for diagnosis and treatment of disorders these systems.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 48

The translation product of this gene shares sequence homology with human polymeric immunoglobulin receptor (accession No.X73079) which is thought to be important in antibody recognition and immune defenses. In one embodiment, polypeptides of the invention comprise the sequence GWYWCG (SEQ ID NO:285). Polynucleotides encoding these polypeptides are also encompassed by the invention.

This gene is expressed primarily in placenta and to a lesser extent in corpus callosum and fetal liver and spleen.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are

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not limited to, disorders of the immune system, e.g. autoimmune diseases and immunodeficiency. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., placenta, liver, and spleen, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:172 as residues: Tyr-37 to Cys-49, Gly-51 to Tyr-56, Lys-88 to Trp-93, Leu-130 to Glu-136.

The tissue distribution and homology to human polymeric immunoglobulin receptor indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of immune disorders, e.g. autoimmune diseases and immunodeficiencies.

FEATURES OF PROTEIN ENCODED BY GENE NO: 49

This gene is expressed in thymus.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune disorder. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., thymus and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of immune disorders, e.g. autoimmunity and immunodeficiency.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 50

Preferred polypeptide encoded by this gene comprise the following amino acid sequence: MKVGARIRVKMSVNKAHPVVSTHWRWPAEWPQMFLHLAQEPRTE VKSRPLGLAGFIRQDSKTRKPLEQETIMSAADTALWPYGHGNREHQENELQKY LQYKDMHLLDSGQSLGHTHTLQGSHNLTALNI (SEQ ID NO:286). Polynucleotides encoding this polypeptide are also provided as are complementary polynucleotides thereto.

This gene is expressed primarily in adrenal gland, pituitary, T helper cells, and breast cells and to a lesser extent in a wide variety of tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of the some diseases and conditions. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and endocrine systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., adrenal gland, pituitary, T-cells and other blood cells, and mammary tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:174 as residues: Gln-39 to Ser-47, Arg-57 to Glu-67, Tyr-82 to Gln-95.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of a wide range of disorders, such as immune and endocrine disorders.

30 FEATURES OF PROTEIN ENCODED BY GENE NO: 51

The translation product of this gene shares sequence homology with human Sop2p-like protein which is important in cytoskeleton structure. In one embodiment, polypeptides of the invention comprise the sequence SLHKNSVSQISVLSGGKAKCS QFCTTGMDGGMSIWDVKSLESALKDLKI (SEQ ID NO:287). Polynucleotides encoding this polypeptide are also encompassed by the invention. This gene maps to chromosome 7. Therefore, polynucleotides of the invention can be used in linkage

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analysis as a marker for chromosome 7.

This gene is expressed primarily in immune and hematopoietic tissues/cells and to a lesser extent in other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immunological and hematopoietic disorders and inflammation. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and hematopoietic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., immune and hematopoietic tissue/cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:175 as residues: Lys-49 to Gln-54, Ala-61 to Arg-66, Lys-82 to Lys-87, Glu-126 to Val-133, His-136 to Ile-141, Glu-175 to Ser-187, Asp-286 to Leu-296, Ala-298 to Ser-310.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of immunological, hematopoietic, and inflammatory disorders, e.g, immunodeficiency, autoimmunity, inflammation.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 52

The translation product of this gene shares sequence homology with *Caenorhabditis elegans* R53.5 gene encoding a putative secreted protein without known function.

This gene is expressed primarily in endothelial cells, brain and several highly vascularized, and tumor tissues and to a lesser extent in other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, aberrant angiogensis and tumorigenesis. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes

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for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the vascular and brain system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., endothelial cells, brain and other tissue of the nervous system, and vascular tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:176 as residues: Thr-43 to Asn-60, Thr-106 to Phe-115, Asp-122 to Arg-133, Arg-186 to Asp-192, Leu-211 to Lys-216.

The tissue distribution and homology to a *C. elegans* secreted protein indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis or treatment of disorders in vascular or brain system, e.g. aberrant angiogenesis, ischemia, neurodegeneration, etc.

FEATURES OF PROTEIN ENCODED BY GENE NO: 53

In one embodiment, polypeptides of the invention comprise the sequence EASKSSHAGLDLFSVAACHRF (SEQ ID NO:288). Polynucleotides encoding this polypeptide are also encompassed by the invention.

This gene is expressed primarily in T-cells and to a lesser extent in brain.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, lymphocytic disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the lymphoid system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:177 as residues: Pro-3 to Thr-8, Arg-37 to Asp-46.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis, treatment, and cure of lymphocytic disorders.

5 FEATURES OF PROTEIN ENCODED BY GENE NO: 54

The translation product of this gene shares sequence homology with secreted cartilage matrix protein, a major component of the extracellular matrix of nonarticular cartilage which is thought to be important in cartilage structure. In specific embodiments, polypeptides of the invention comprise the sequence: RCKKCTEGPI DLVFVIDGSKSLGEENFEVVKQF (SEQ ID NO:297); VTGIIDSLTISPKAARVGL LQYSTQVH (SEQ ID NO:290); TEFTLRNFNSAKDMKKAVAHMKYM (SEQ ID NO:291); GKGSMTGLALKHMFERSFTQGEGARPF (SEQ ID NO:292); STRVP RAAIVFTDGRAQDDVSEWASKAKANGITMYAVGVGKAIE (SEQ ID NO:293); EELQEIASEPTNKHLFYAEDFSTMDEISEKLKKGICEALEDS (SEQ ID NO:294); TQRLEEMTQRM (SEQ ID NO:295); PQGCPEQPLH (SEQ ID NO:296); and/or YMGKGSMTGLALKHMFERSFT (SEQ ID NO:289). Polynucleotides encoding these polypeptides are also encompassed by the invention.

This gene is expressed primarily in placenta, infant brain, prostate, fetal lung and to a lesser extent in endometrium and fetal tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, abnormal placenta and pregnancy, disorder and injury in brain, prostate, and vasculature. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the reproduction, neuronal, and vascular systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., placenta, brain and other tissue of the nervous system, prostate, lung and endometrium, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to cartilage matrix protein indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis,

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treatment, and cure of abnormalities in placenta and pregnancy, disorder and injury in brain, prostate, and vasculature.

FEATURES OF PROTEIN ENCODED BY GENE NO: 55

The translation product of this gene is the human ortholog of bovine and hamster CII-3, a succinate-ubiquinone oxidoreductase complex II membrane-intrinsic subunit, which is thought to be important in mitochondrial electron transport chain during metabolism. In specific embodiments, the polypeptides of the invention compriseMAALLLRHVGRHCLRAHFSPQLCIRNAVPLGTTAKEEMERFWNKNIG SNRPLSPHITIYS (SEQ ID NO:298); VFPLMYHTWNGIRHLMWDLGKGLKIPQL YQSG (SEQ ID NO:299); MAALLLRHVGRHCLRAH (SEQ ID NO:300); VKSLCL GPALIHTAKFAL (SEQ ID NO:301); VFPLMYHTWNGIRHLMWDLGKGL (SEQ ID NO:302).

This gene is expressed in 8-week old early stage human.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, metabolism disorder. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the [insert system where a related disease state is likely, e.g., immune], expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis, treatment, and cure of metabolism disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 56

This gene is expressed primarily in umbilical vein endothelial cells, human ovarian tumor cells, human meningima cells, and human Jurkat membrane bound polysomes. In specific embodiments, polypeptides of the invention comprise the amino acid sequence: RVWDVRPFAPKERCVKIFQGNV (SEQ ID NO:303); HNFEKNLL

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RCSWSPDGSKIAAGSADRFVYV (SEQ ID NO:304); and/or WDTTSRRILYKLPG HAGSINEVAFHPDEPI (SEQ ID NO:305). Polynucleotides encoding these polypeptides are also encompassed by the invention.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, inflammation, immune and cardiovascular disorders and urogenital neoplasias. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, neurological, urogenital, reproductive system and vascular systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., blood cells, cells, endothelial cells, ovary and other reproductive tissue, meningima, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:143 as residues: Phe-71 to Arg-76, Pro-82 to His-87, Glu-103 to Ala-111.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the diagnosis and treatment of immune disorders including: leukemias, lymphomas, auto-immune, immuno-supressive (e.g. transplantation) and immunodeficiencies (e.g. AIDS) and hematopoietic disorders. In addition, expression in ovarian tumor cells suggests that polynucleotides and polypeptides corresponding to this gene are useful for study, diagnosis, and treatment of ovarian tumors, and other tumors and neoplasias. Further, endothelial cell expression suggests a role in cadiovascular or respiratory/pulmonary disorders or infections (athsma, pulmonary edema, pneumonia).

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FEATURES OF PROTEIN ENCODED BY GENE NO: 57

The translation product of this gene shares sequence homology with type I collagen. In specific embodiments, the polypeptides of the invention comprise the sequence: GRIPAPAPSVPAGPDSR (SEQ ID NO:309); VRGRTVLRPGLDAEPE LSPE (SEQ ID NO:306); EQRVLERKLKKERKKEERQ (SEQ ID NO:307); ARRSG

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AELAWDYLCRWAQKHKNWRFQKTRQTWLLLHMYDSDKVPDEHFSTLLAYLE GLQGR (SEQ ID NO:255); and/or RLREAGLVAQHPP (SEQ ID NO:308). Polynucleotides encoding these polypeptides are also encompassed by the invention.

This gene is expressed primarily in epididymus, prostate cell line (LNCAP), and pituitary gland; and to a lesser extent in many other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, abnormalities of the epididymus, prostate (especially prostate cancer), and pituitary gland. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the male reproductive system and neuroendocrine system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., epididymus and other reproductive tissue, prostate, and pituitary gland, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to type I collagen, indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of abnormalities of the epididymus, prostate (especially prostate cancer), and pituitary gland.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 58

This gene is expressed primarily in the frontal cortex of the brain from a schizophrenic individual.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, schizophrenia. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the nervous system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., brain and other tissue of

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the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of schizophrenia.

FEATURES OF PROTEIN ENCODED BY GENE NO: 59

The polypeptide encoded by Gene 59 is homologous to human surface 4 integral membrane protein. In specific embodiments, the polypeptides of the invention comprise the sequence: TGCVLVLSRNFVQYACFGLFGIIALQTIAYSILWDLKF LMRN (SEQ ID NO:310); SRSEGKSMFAGVPTMRESSPKQYMQLGGRVLLV LMFMTLLHFDASFFSIVQNIVG (SEQ IDNO:311); GTAEDFADQFLRVTKQYLP HVARLCLISTFLEDGIRMFQWSEQRDYIDTTWNCGYLLAS (SEQ ID NO:312); LMRNESRS (SEQ ID NO:314); ASFLLSRTSWGTA (SEQ ID NO:315); and/or ASFLLSRTSWGTALMIL (SEQ ID NO:313). Polynucleotides encoding these polypeptides are also encompassed by the invention.

This gene is expressed primarily in Hodgkin's lymphoma and lung; and to a lesser extent in many other human tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, Hodgkin's lymphoma, tumors or other abnormalities of the lung. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and respiratory systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., lymphoid tissue, and pulmonary tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO: 183 as residues: Met-20 to Trp-27.

The tissue distribution indicates that polynucleotides and polypeptides

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corresponding to this gene are useful for diagnosis and treatment of Hodgkin's lymphoma, tumors or other abnormalities of the lung.

FEATURES OF PROTEIN ENCODED BY GENE NO: 60

This gene is expressed primarily in bone cancer and stomach cancer, and to a lesser extent in many other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, bone cancer and stomach cancer. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the bone, and the stomach, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., bone, and stomach, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of bone cancer and stomach cancer and possibly other cancers.

FEATURES OF PROTEIN ENCODED BY GENE NO: 61

This gene is expressed primarily in epididymus, and lymph node of breast cancer, and to a lesser extent in many other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, abnormalities of the epididymus, and breast cancer. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the epididymus and breast, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., epididymus and other reproductive tissue, lymphoid tissue, and mammary tissue, and cancerous and wounded tissues) or bodily fluids

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(e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:185 as residues: Arg-57 to Ser-65.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of abnormalities of the epididymus, and breast cancer.

FEATURES OF PROTEIN ENCODED BY GENE NO: 62

The translation product of this gene appears to be the human homolog of bovine NADH dehydrogenase which is thought to be important in cellular metabolism. In specific embodiments, the polypeptides of the invention comprise the amino acid sequence: SMSALTRLASFARVGGRLFRSGCARTAGDGGVRHAGGGVHIEPRY RQFPQLTRSQVFQSEFFSGLMWFWILWRFWHDSEEVLGHFPYPDPSQWTDEEL GIPPDDED (SEQ ID NO:323), or fragments thereof. Polynucleotides encoding this polypeptide are also encompassed by the invention.

This gene is expressed in larynx tumor, lymph node, brain amygdala, human cardiomyopathy, and retina.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, diseases affecting cellular metabolism. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the nervous system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., larynx, lymphoid tissue, brain and other tissue of the nervous system, heart and cardiovascular tissue, and retina, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:208 as residues: Pro-27 to Gln-32, Arg-42 to Glu-51.

The tissue distribution and homology to NADH dehydrogenase indicates that

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polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of diseases involving cellular metabolism.

FEATURES OF PROTEIN ENCODED BY GENE NO: 63

This gene is expressed primarily in amygdala, and to a lesser extent in many other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, abnormalities of the amygdala. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the amygdala, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., amygdala, and lymphoid tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:187 as residues: Gln-17 to Glu-29, Pro-41 to Phe-46, Ser-59 to Ile-70, Thr-97 to Leu-105.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of abnormalities of amygdala.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 64

This gene is expressed primarily in female bladder, and to a lesser extent in chronic synovitis and hemangiopericytoma.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, bladder cancer. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the urinary tract, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., bladder, synovial tissue, and

vascular tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:188 as residues: Pro-2 to Gln-7, Pro-27 to Phe-34.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatments of defects of the urinary tract, especially bladder cancer.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 65

This gene is expressed primarily in fetal spleen, and to a lesser extent in hemangiopericytoma, thymus, and synovial sarcoma.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, defects of immune of hematopoietic systems. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune of hematopoietic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., spleen, vascular tissue, thymus, blood cells, and synovial tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The protein product of this gene is useful for treatment of defects of the immune or hematopoietic systems, because of the gene's expression in thymus and spleen.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 66

This gene is expressed primarily in human pituitary and to a lesser extent in placenta and fetal lung.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are

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not limited to, endocrine growth disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the endocrine system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., pituitary and other endocrine tissue, placenta, and pulmonary tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:190 as residues: Val-38 to Asn-44, Gly-53 to Ser-65.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment of growth disorders related to pituitary dysfunction.

FEATURES OF PROTEIN ENCODED BY GENE NO: 67

The translation product of this gene shares sequence homology with a *Caenorhabditis elegans* gene of unknown function. In specific embodiments, the polypeptides of the invention comprise the sequence: DPRRPNKVLRYKPPPSE CNPALDDPTP (SEQ ID NO:317); DYMNLLGMIFSMCGLMLKLKWCAWVA VYCS (SEQ ID NO:318); FISFANSRSSEDTKQMMSSF (SEQ ID NO:316); and/or MLSISAVVMSYLQNPQPMTPPW (SEQ ID NO:319). Polynucleotides encoding these polypeptides are also encompassed by the invention.

This gene is expressed primarily in primary breast cancer and lymph node breast cancer and to a lesser extent in adult brain, lung cancer, colon cancer, epithelioid sarcoma, and Caco-2 cell line.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cancer. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the cancer and tumor tissues, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., mammary tissue, lymphoid tissue, brain and other tissue of the nervous system, lung, colon, and

epithelium, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:191 as residues: Asn-34 to Lys-42.

The tissue distribution in a variety of cancer tissues indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of a variety of cancer and tumor types.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 68

The translation product of this gene shares sequence homology with steroid membrane binding protein. The translation product of this gene has recently been published as progesterone binding protein. See Genbank AJ002030. Preferred polypeptides encoded by this gene comprise the following amino acid sequence: AAGDGDVKLGTLGSGSESSNDGGSESPGDAGAAAXGGGWAAAALALLTG GGE (SEQ ID NO:320).

This gene is expressed primarily in breast, and to a lesser extent in placenta and fetal tissue.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, breast cancer or developmental disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders 25 of the above tissues or cells, particularly of breast or fetal tissues, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., mammary tissue, placenta, and fetal tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the 30 standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:192 as residues: Pro-43 to Asp-49, Gln-54 to Pro-64, Asp-110 to Asp-118, Lys-138 to Tyr-143, Pro-150 to Asp-170.

The tissue distribution and homology to steroid membrane binding protein and to progesterone binding protein indicates that the protein products of this gene are

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useful for treatment of breast cancers, especially those caused by estrogen and progesterone binding.

FEATURES OF PROTEIN ENCODED BY GENE NO: 69

Preferred polypeptides encoded by this gene comprise the following amino acid sequence: AADNYGIPRACRNSARSYGAAWLLLXPAGSSRVEPTQDISISDQLGG QDVPVFRNLSLLVVGVGAVFSLLFHLGTRERRPHAXEPGEHTPLLAPATAQPL LLWKHWLREXAFYQVGILYMTTRLIVNLSQTYMAMYLTYSLHLPKKFIATIPLV MYLSGFLSSFLMKPINKCIGRN (SEQ ID NO:321).

This gene is expressed primarily in macrophage (GM-CSF treated), and to a lesser extent in monocytes and dendritic cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, inflammation and infection. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., macrophages and other blood cells, and dendritic cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that the protein products of this gene are useful for treatment of infection or inflammation or other events or defects involving the immune system.

FEATURES OF PROTEIN ENCODED BY GENE NO: 70

This gene is expressed primarily in adult brain and to a lesser extent in thyroid, 12 week old early stage human, and stromal cell TF274.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, neurological or neuro-endocrine diseases. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes

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for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the central nervous or endocrine systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., brain and other tissue of the nervous system, thyroid, and stromal cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:194 as residues: Pro-65 to Cys-71.

The tissue distribution indicates that the protein products of this gene are useful for treatment and diagnosis of neurological diseases or metabolic conditions involving the neuro-endocrine system.

FEATURES OF PROTEIN ENCODED BY GENE NO: 71

This gene is expressed in T-cell helper and to a lesser extent in adult brain and adult testes.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune disorders, meningitis or reproductive problems. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, neural and reproductive systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, brain and other tissue of the nervous system, testes and other reproductive tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:195 as residues: Val-18 to Tyr-24, Ala-89 to Asp-99, Asp-104 to Ala-117, Leu-121 to Pro-136.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis immune and

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reproductive disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 72

The translated polypeptide of this contig has a high degree of identity with the Ob Receptor-Associated Protein deposited as GenBank Accession No. 2266638. No function has been determined for the Ob Receptor-Associated Protein, however it is expressed upon stimulation of the Ob Receptor by Leptin.

This gene is expressed in T-cells and to a lesser extent in endothelial and bone marrow cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, acute lymphoblastic leukemia, hematapoetic disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and hematapoetic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, endothelial cells, and bone marrow, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:196 as residues: Ser-61 to Trp-70.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of leukemia and other disorders of the primary immune system. In addition, since this gene appears to be related to the Ob Receptor-Related Protein, it is likely that this polypeptide is also involved in the Ob/Leptin signal transduction cascade. As a result, this protein may be of use in the molecular diagnosis and therapeutic intervention of obesity and related disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 73

The translation product of this contig has homology with furin, a protein thought to be a key endopeptidase in the constitutive secretory pathway. The identification and initial characterization of Furin was reported by Takahasi and

colleagues (Biochem Biophys Res Commun 1993 Sep 15;195(2):1019-1026).

This gene is expressed in neutrophils.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, diseases of the immune system such as allergies, wound healing and antigen recognition. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., neutrophils and other blood cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment of allergies or other immune disorders since neutrophils are an important part of an allergic response. Further, since this protein appears to be related to Furin, it can be used diagnostically and therapeutically to treat secretory protein processing disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 74

This gene is expressed in the frontal cortex.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, of the motor activity and sensory functions that involve the central nervous system. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the central nervous system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene

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expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the detection and treatment of neural disorders that affect cognitive functions.

FEATURES OF PROTEIN ENCODED BY GENE NO: 75

The translation product of this gene shares sequence homology with inorganic pyrophophatase which is thought to be important in the catalysis the hydrolysis of diphosphate bonds, chiefly in nucleoside di- and triphosphates and essential enzymes that are important for controlling the cellular levels of inorganic pyrophosphate (PPi). The bovine homolog of this gene has been identified by Yang and Wensel (J. Biol. Chem. 267:24641-24647 (1992)).

This gene is expressed in osteoclastoma cells and to a lesser extent in epithelial cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, osteoporosis and other bone weakening diseases. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the skeletal system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., bone, and epithelial cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:199 as residues: Lys-22 to Tyr-28, Asp-64 to Lys-77, Pro-86 to Ile-91, Gln-99 to Pro-119, Tyr-169 to Asp-174, Lys-176 to Gly-181, Trp-189 to Asn-202, Lys-233 to Gly-239, Ser-250 to Asp-257.

The tissue distribution and homology to inorganic pyrophophatase indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of osteoporosis through the removal of bone by demineralization.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 76

The translation product of this gene shares exact sequence homology with ATP sulfurylase/APS kinase (GenBank Accession No. 2673862) which is thought to be important in biosynthesis of the activated sulfate donor, adenosine 3'-phosphate 5'-phosphosulfate, involves the sequential action of two enzyme activities: ATP sulfurylase, which catalyzes the formation of adenosine 5'-phosphosulfate (APS) from ATP and free sulfate, and APS kinase, which subsequently phosphorylates APS to produce adenosine 3'-phosphate 5'-phosphosulfate.

This gene is expressed in osteoclastoma cells and to a lesser extent in developmental tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, antibiotic resistant bacterial infections, osteoarthritis and other auto immune diseases. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune or skeletal structure expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., bone, and developmental tissues, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:200 as residues: Asn-15 to Trp-20, Ser-36 to Gly-41, Pro-103 to Val-110, Pro-134 to Arg-143, Leu-173 to Arg-178, Ser-190 to Ala-197, His-314 to Arg-319, Arg-354 to Asn-362, Asp-391 to Arg-397, Glu-402 to Asp-409, Asp-434 to Leu-439, Glu-441 to Arg-446, Gly-455 to Asp-462, Pro-528 to His-541, Asn-566 to Arg-571, Tyr-574 to Glu-581, Thr-589 to Glu-603.

The tissue distribution and homology to ATP sulfurylase/APS kinase indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment or detection of autoimmune diseases.

FEATURES OF PROTEIN ENCODED BY GENE NO: 77

This polypeptide is identical to the SLP-76-associated protein reported by Musci and colleagues (J. Biol. Chem. 272 (18), 11674-11677 (1997)) and to the FYB protein

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reported by da Silva and coworkers (Proc. Natl. Acad. Sci. U.S.A. (1997) In press). These proteins have been reported to be novel T-cell Proteins which bind FYN and SLP-76 and regulate IL-2 production. Preferred polypeptides encoded by this gene comprise the following amino acid sequence: RITDNPEGKWLGRTARGSYGYIK TTAVEIXYDSLKLKKDSLGAPSRPIEDDQEVYDDVAEQDDISSHSQSGSGGIFPP PPDDDIYDGIEEEDADDGFPAPPKQLDMGDEVYDDVDTSDFPVSSAEMSQGTNV GKAKTEEKDLKKLKKQXKEXKDFRKKFKYDGEIRVLYSTKVTTSITSKKWGT RDLQVKPGESLEVIQTTDDTKVLCRNEEGKYGYVLRSYLADNDGEIYDDIADGC IYDND (SEQ ID NO:322).

This gene is expressed in CD34 positive cells (hematopoietic progenitor cells) and to a lesser extent in adult spleen derived from a chronic lymphocytic leukemia patient.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are 15 not limited to, chronic lymphocytic leukemia; hematopoietic disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and hematopoietic systems, expression of this gene at significantly higher or lower levels 20 may be routinely detected in certain tissues (e.g., T-cells and other blood cells, bone marrow, hematopoietic cells, and spleen, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the 25 standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Further, nucleic acids and polypeptides of the present invention are useful both diagnostically and therapeutically in the intervention of immune and other disorders in which the ability to alter IL-2 expression is desired. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:201 as residues: Ala-17 to Lys-37, Val-39 to Ser-45, Lys-59 to His-70, 30 Arg-90 to Leu-95, Lys-97 to Lys-107, Ser-117 to Leu-124, Phe-133 to Ser-138, Trp-146 to Leu-167, Pro-175 to Asn-185, Lys-190 to Ser-211, Pro-213 to Ser-222, His-230 to Pro-235, Pro-240 to Pro-246, Pro-253 to Gly-261, Leu-271 to Leu-303, Leu-305 to Leu-326, Lys-343 to Leu-349, Thr-363 to Leu-371, Arg-373 to Tyr-381, Tyr-391 to Leu-401, Pro-404 to Val-414, Ser-426 to Ser-432, Ile-448 to Ser-457, Gln-462 35 to Trp-468, Lys-477 to Ser-501, Asp-518 to Ser-523, Ala-541 to Gln-554.

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The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment of a variety of hematopoietic disorders. The noted expression of this gene in the hematopoietic progenitor cell compartment - as determined by its expression on CD34 positive hematopoietic stem and progenitor cells - indicates that it plays a critical role in the expansion or proliferation of hematopoietic stem/progenitor cells, as well as in the differentiation of the various blood cell lineages. Thus it could be useful in the reconstitution of the hematopoietic system of patients with leukemias and other hematopoietic diseases.

10 FEATURES OF PROTEIN ENCODED BY GENE NO: 78

This gene is homologous to heparin cofactor II (HCII) which is a 66-kDa plasma glycoprotein that inhibits thrombin rapidly in the presence of dermatan sulfate or heparin.

This gene is expressed in apoptotic and anergic T-cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, thrombopienia T-cell lymphomas; Hodgkin's lymphoma. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system most notably the T-cell compartment, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, and lymphoid tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The homology to heparin cofactor II (HCII) and the tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of hematopoietic disorders particularly in thrombopoesis, most notably of the T-cell compartment. This could include immune modulation, inflammation, immune surveillance, graft rejection, and autoimmunity.

35 FEATURES OF PROTEIN ENCODED BY GENE NO: 79

The translation product of this gene shares sequence homology with a mouse

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protein believed to represent an integral membrane protein.

This gene is expressed in fetal cochlea and epididymus and to a lesser extent in adult spleen and osteoclastoma.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, osteoclastoma; disorders of the inner ear; male fertility disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the inner ear; male reproductive tract; bone; and immune systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., cochlea, epididymus and other reproductive tissue, spleen, and bone, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:203 as residues: Lys-13 to Gly-23, Cys-38 to Asp-43, Gly-48 to Trp-53, Cys-223 to Ile-237, Ile-240 to Ser-246.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment of hearing and fertility disorders. Likewise, it may have a role in the modulation of immune function and in the treatment of osteoporosis.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 80

The translation product of this gene shares sequence homology with reticulocalbin which is thought to be important in the binding of calcium, particularly within the endoplasmic reticulum.

This gene is expressed in endothelial cells and stromal cells and to a lesser extent in osteoblasts, osteoclasts, and T-cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, osteoperosis; osteoclastomas; T-cell lymphomas; Hodgkin's disease. Similarly, polypeptides and antibodies directed to these polypeptides are useful in

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providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the vasculature, bone, and immune systems - particularly the T-cell compartments, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., endothelial cells, stromal cells, bone, T-cells and other blood cells, and lymphoid tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:204 as residues: Lys-20 to Arg-27, Pro-32 to Asp-48, Leu-64 to Arg-72, Asp-108 to Lys-114, Glu-128 to Thr-133, Asp-139 to Phe-147, Thr-196 to Ala-204, Tyr-218 to Glu-228, Val-230 to Gln-236, Arg-241 to Lys-255, Glu-276 to Lys-287.

The tissue distribution and homology to reticulocalbin indicates that polynucleotides and polypeptides corresponding to this gene are useful for the diagnosis and treatment of bone disorders such as osteoporosis; the diagnosis and treatment of T-cell lymphomas and Hodgkin's lymphoma; and the treatment of diseases and defects of the vasculature, such as vascular leak syndrome and aberrant angiogenesis that accompanies tumor growth.

FEATURES OF PROTEIN ENCODED BY GENE NO: 81

The translation product of this gene shares sequence homology with a family of peptide transport genes - particularly the AtPTR2-B gene from *Arabidopsis* - which are thought to be important in the uptake of small peptides.

This gene is expressed in a number of fetal tissues, most notably lung, brain, cochlea, and liver/spleen, and to a lesser extent in osteoclastoma and endometrial tumors.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, osteoclastoma; endometrial tumors; cancer; leukemias. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the bone and endometrium, expression of this gene at significantly higher or lower levels may be

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routinely detected in certain tissues (e.g., fetal tissue, pulmonary tissue, bone, brain and other tissue of the nervous system, cochlea, liver, and spleen, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:205 as residues: Lys-186 to Asn-199, Pro-202 to Ala-207.

The tissue distribution and homology to peptide transport genes indicates that polynucleotides and polypeptides corresponding to this gene are useful for the control of cell proliferation, owing to its strong expression in fetal tissues undergoing active cell division, as well as its expression in a variety of tumors or cancers of adult tissues. Potentially, it may regulate the uptake of peptides that stimulate cell proliferation. This gene product may also be useful in stimulating the uptake of a variety of peptide-based drug compounds.

FEATURES OF PROTEIN ENCODED BY GENE NO: 82

This gene is expressed in fetal liver and spleen and to a lesser extent in endothelial cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cancer and tumors of a hematopoietic and/or endothelial cell origin; leukemias. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system and/or vasculature, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., liver, spleen, endothelial cells, vascular tissue, and tissue and cells of the immune system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:206 as residues: Met-1 to Asp-9, Arg-66 to Gly-76, Asp-164 to Arg-171.

The tissue distribution indicates that polynucleotides and polypeptides

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corresponding to this gene are useful for the treatment of disorders of the immune system. Expression of this gene product in both fetal liver/spleen and endothelial cells indicates that it may be expressed in the hemangioblast, the progenitor cell for both the immune system and the vasculature. Thus, it is most likely expressed in hematopoietic stem cells, and may be useful for the expansion of hematopoietic stem and progenitor cells in conjunction with cancer treatment for a variety of leukemias.

FEATURES OF PROTEIN ENCODED BY GENE NO: 84

The translation product of this gene shares sequence homology with NADH dehydrogenase which is thought to be important in cellular metabolism.

This gene is expressed in fetal dura mater and to a lesser extent in T-cells and hypothalamus.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, diseases affecting cellular metabolism. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the nervous system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., fetal tissue, T-cells and other blood cells, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:208 as residues: Pro-27 to Gln-32, Arg-42 to Glu-51.

The tissue distribution and homology to NADH dehydrogenase indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of diseases involving cellular metabolism.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 85

The translation product of this gene shares sequence homology with I-TRAF, a novel TNF receptor associated factor (TRAF)-interacting protein that regulates TNF receptor-mediated signal transduction. This protein is thought to be important in regulating the cellular response to tumor necrosis factor (TNF), which is an important mediator of inflammation.

This gene is expressed in endothelial cells and to a lesser extent in glioblastoma and osteoblastoma.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, inflammation; glioblastoma and osteoblastoma. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., endothelial cells, bone, and glial cells and tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:209 as residues: Glu-15 to Thr-22, Glu-46 to Leu-62, Arg-103 to Glu-119, Gln-127 to Glu-132, Asn-152 to Trp-158, Gln-191 to Gln-210, Glu-264 to Thr-271, Tyr-282 to Leu-288, Trp-319 to Thr-331, Glu-335 to Ser-348, Ser-353 to Ser-358, Asp-382 to Asn-392.

The tissue distribution and homology to I-TRAF indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of inflammatory diseases, including rheumatoid arthritis, sepsis, inflammatory bowel disease, and psoriasis, particularly where tumor necrosis factor is known to be involved.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 86

This gene has homology with a candidate gene involved in X-linked Retinopathy reported by Wong and colleagues (Genomics 15:467-471 (1993)).

This gene is expressed in a T-cell line.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, inflammation and autoimmune diseases; T-cell lymphoma. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of inflammatory disorders such as sepsis, inflammatory bowel disease, psoriasis, and rheumatoid arthritis as well as autoimmune disease such as lupus. It could also be useful in immune modulation and in the process of immune surveillance. The present invention can be used diagnostically and therapeutically to treat X-linked Retinopathy.

25 FEATURES OF PROTEIN ENCODED BY GENE NO: 87

This gene is expressed in human brain tissue.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, brain disorders; neurodegenerative disorders; tumors of a brain origin. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the central nervous system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma,

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urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:211 as residues: Cys-32 to Tyr-38.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of CNS disorders such as epilepsy, paranoia, depression, Alzheimer's disease, and schizophrenia. It could be useful in the survival and/or proliferation of neurons and could effect neuronal regeneration.

Last AA of ORF	30	44	69	-	38	22	601
Last AA First AA of of Sig Secreted Pep Portion (27	45	26	25	23	21
Last AA of Sig Pep		26	44	25	24	22	20
irst AA of of Sig Pep	I	-	-	_	_	_	-
AA SEQ D NO: Y	125	126	212	213	127	128	129
of AA F First SEQ AA of ID Signal NO:	353	128	170	413	66	006	103
NT of tart	353	128	170	413	66	006	103
3' NT of Clone Seq.	1607	1786	1487	1637	1212	2061	733
Total Clone Clone S S Seq. Seq. Seq.	247	87	79	394		882	01
Total NT Seq.	1679	1830	1487	1653	1212	2061	1412
X SEQ	Ξ	12	86	66	13	4	15
Vector	Uni-ZAP XR	Uni-ZAP XR	pBluescript	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR
ATCC Deposit Nr and Date	97923 03/07/97 209071	97923 03/07/97 209071	86/61/EU	209641	97923 97923 03/07/97 209071	97923 03/07/97 209071	03/22/97 97923 03/07/97 209071 05/22/97
cDNA Clone ID	HAGEW82	HAGFY16	HBMCF37	HFLQB16	HALAA60	HAPBL78	HASAV70
Gene No.		2	2	2	3	4	2

Last AA of ORF	62	29	52	56	215	48
First AA of Secreted Portion	8		24	18	61	27
First Last AA AA of of Sig Sig Pep Pep	17		23	17	<u>∞</u>	26
First AA of Sig Pep		_	-	-	_	_
¥ŠeŠ;≻	130	131	132	133	134	135
5' NT of First AA of Signal Pep	538	181	98	192	401	793
5' NT of AA F of AA of E's NT First SEQ / Of AA of D Start Signal NO: Star	538	181	98	192	401	793
S' NT 3' NT of of Olone Clone Seq. Seq.	088	683	1007	1393	1070	2011
s, NT of Clone Seq.	276	_	98	132	277	614
Total NT Seq.	1052	683	1054	1393	1215	2042
SEQ NO: NO:	16	17	<u>8</u>	61	20	21
Vector	Uni-ZAP XR					
ATCC Deposit Nr and Date	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97
cDNA Clone ID	HBNAF22	HBNBL77	HCDDR90		HCEMU42	HCENE16
Gene No.	9	7	8	6	01	11

Last AA of ORF	/9	21	539	08	56	84	200
ed on	24	30	31	23	27	3/	28
	23	67.	30	22	26	36	27
First AA of Sig Pep			_	-	_ -		-
· · · · · · · · · · · · · · · · · · ·	136	137	138	214	139	215	140
al of t	69	68	808	515	961	295	10
5' NT of Start Codon	69	89	808	515	961	295	70
3' NT of Clone Seq.	1872	289	3532	1115	907	734	717
5' NT 3' NT of Clone Clone Seq.	21	_	2821	435	171	25	_
Total NT Seq.	1872	586	3533	1145	1148	734	717
SEQ NÖ:	22	23	24	100	25	101	26
Vector	Uni-ZAP XR	ZAP Express	Uni-ZAP XR	Uni-ZAP XR	pBluescript	Uni-ZAP XR	Uni-ZAP XR
ATCC Deposit Nr and Date	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/27/97	97923 03/07/97 209071	209179 209179 07/24/97	97923 03/07/97 209071 05/22/97	97923 97923 03/07/97 209071 05/22/97
cDNA Clone ID	HMSJJ74	HCUBF15	HE2DE47	HE2DE47	HKMLH01	HE6DG34	HE9DG49
Gene No.	12	13	14	14	15	15	16

Last AA of ORF	202	215	185	101	=	61
irst AA of Secreted Portion	29	23	26	43	31	
Last AA of Sig Pep	28	22	25	42	30	
First AA of Sig Pep		-	_		_	
¥ŠEŞ ∀ÖÐÖ;≻	216	141	217	142	143	144
of AA First Last AA of D of of Signal NO: Sig Sig S A Pep Pep	78	38	149	128	294	496
of of Start	78	38	149	128	294	496
S' NT 3' NT of of Clone Clone Seq. Seq.	713	1099	1080	941	756	2093
S' NT of Clone Seq.	17	_	_	171	79	408
Tota NT Seq	713	6601	1080	941	756	2100
NT SEQ ID NO:	102	27	103	28	29	30
Vector	Uni-ZAP XR					
ATCC Deposit Nr and Date	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97
	HE9DG49	HELBA06	HELBA06		HELBW38	HETHN28
Gene No.	16	17	17	18	19	20

		//					
1 ' 0	67.	0,	_	38	130	3.1	
First AA of Secreted Portion	QC	67		17	27	22	
Last AA of Sig Pep	oc C	07		16	26	21	
First AA of Sig Pep		T		_		_	_
SEQ YÖ: BÖ	145	140	147	148	149	150	151
	567	17	210	242	178	144	1104
5' NT of Start Codon	567	21	210	242	178	144	1104
	1392	409	1322	710	1161	938	1581
	475	-	1		110	_	974
Total NT Seq.	1448	456	1326	710	1188	956	1603
XÖ: BÖ	31	32	33	34	35	36	37
Vector	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR
ATCC Deposit Nr and Date	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/27/97	97923 03/07/97 209071	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97924 03/07/97
cDNA Clone ID	HFCDK17	HFEAF41	HFKFL13	HFSBG13	HFTBE43	HFTDJ36	HKTAC77
Gene No.	21	22	23	24	25	26	27

Last AA of ORF	7	29	25	194	8	30	68	68	88	173	137	47	44
First AA of Secreted Portion		33		33	19	31	20	23	19	21	21	28	28
Last AA of Sig Pep		32		32	18	30	61	22	81	20	20	27	27
First AA of Sig Pep	_	_			_	1	_	_			-	-	-
SEQ YÖ: BÖ:	152	153	154	155	156	157	158	218	159	091	219	220	191
5' NT of First AA of Signal Pep	209	119	581	126	43	171	55	58	17	15	72	54	269
5' NT of Start Codon		611	581	126	43	171	55	58	17	15	72	54	569
S' NT 3' NT of of Clone Clone Seq. Seq.	1901	629	1793	1123	875	843	489	489	534	1374	640	1399	596
5' NT of Clone Seq.	55	_	408	13	_	-	3	9	_	_	58	40	
Total NT Seq.	1089	629	1964	1522	875	843	489	489	534	1374	640	1529	969
SEQ BD NO:	38	39	40	41	42	43	44	104	45	46	105	106	47
Vector	pBluescript	pBluescript	Lambda ZAP II	Uni-ZAP XR	Uni-ZAP XR	pSport1	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR				
ATCC Deposit Nr and Date	97924 03/07/97	97924	97924 03/07/97										
cDNA Clone ID	нснѕнз6	HLHSV96	98ОВОТН	HLTBX31	HLTCJ63	HMKAH44	HMQAJ64	HMQAJ64	HOABG65	HODCL36	HODCL36	HODCL36	HODCL50
Gene No.	28	29	30	31	32	33	34	34	35	36	36	36	37

Last AA of ORF	22	69	775	6	519	78	30	=	280	7 4 6	7.7	326	183
First AA of Secreted Portion		81	0.7	75	19	7.7	,	61	31	31		20	24
Last AA of Sig Pep		17	13	31	09	21		8	30	ر م		61	23
First AA of Sig Pep	-		1	1		_	-	_		_	_	_	
AA SEQ ID NO: Y	162	163	104	777	165	222	166	167	168	223	169	170	224
5' NT of First AA of Signal Pep	170	638	99	928	150	239	432	142	25	433	217	57	35
S' NT of Start Codon	170	638	99	928	150	239	432	142	25	433	217	57	35
3' NT of Clone Seq.	822	2020	2432	2435	2340	791	601	337	1141	1166	1148	809	286
5' NT 3' NT of of Clone Clone Seq. Seq.	66	569		849	1627	92	188	_	-	21	63	164	4
Total NT Seq.	851	2020	2432	2435	2340	805	601	359	1141	1166	1560	1507	586
SEQ NÖ:SEQ	48	49		107	51	108	52	53	54	601	55	56	110
Vector	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	pBluescript SK-	pBluescript SK-	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR
ATCC Deposit Nr and Date	97924	97924 03/07/97	97924 03/07/97	97924 03/07/97	97924 03/07/97	97924	97924 03/07/97	97924	97924 03/07/97	97924 03/07/97	97924	97924	97924 03/07/97
cDNA Clone ID	HODCV74	HODCZ16	HTOEU03	HTOEU03	HPBCJ74	HPBCJ74	HPMBU33	HSAUL66	HSIDQ18	HSIDQ18	HSJBB37	HSJBQ79	HSJBQ79
Gene No.	38	39	40	40	41	41	42	43	44	44	45	46	46

Last AA of ORF	89	158	70	122	128	6	371
First AA of Secreted Portion	36	91	20	61	31		2
Last AA of Sig Pep	35	15	61	<u>&</u>	30		_
First AA of Sig Pep	_	_	_		_	-	-
¥ŠEQ ⊀ÖÐĞ	171	172	225	173	174	226	175
S' NT of AA F First SEQ AA of D Signal NO: 9 Pep Y F	83	163	155	115	52	829	114
5' NT of Start Codon	83	163	155	115	52	829	114
S' NT 3' NT of Ol Clone Clone Seq. Seq.	450	1147	1134	777	598	1333	1554
	1		-	-	48	594	443
Total NT Seq.	450	1147	1134	777	1611	1333	1580
SEQ NÖ BÖ	57	28	111	59	09	112	61
Vector	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	pBluescript	pBluescript	Uni-ZAP XR
ATCC Deposit Nr and Date	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	209235 09/04/97
	HTEGA76	HTEJN13	HTEJN13	HTHBL86	HTSF071		HAPNO80
Gené No.	47	48	48	49	50	50	51

. – – 1	137	CI7	54	22	102	47
First AA of Secreted Portion	29	67	33	21	34	39
Last AA of Sig Pep	28	87	32	_20	33	38
First AA of Sig Pep	-			1		_
AA SEQ ID NO: Y	227	176	177	8/1	179	180
Start Signal NO: Start Sep NY Start Signal NO: Start Sign	244	182	<i>L</i> 6	150	231	703
5' NT of Start Codon	244	182	26	150	231	703
3' NT of Clone Seq.	708	1034	361	1638	1303	1011
NT SEQ D Total Clone Clone of NO: NT Seq. Seq. Stat X Seq.	249	501	_	-	35	655
Total NT Seq.	1015	1117	361	1668	1353	1011
SEQ SEQ X	113	62	63	64	65	99
Vector	Uni-ZAP XR	pBluescript	pSport1	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR
ATCC Deposit Nr and Date	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 97958 03/13/97 209072 05/22/97
cDNA Clone ID	HAUCC47	HBMCL41	HCFLD84	НЕ8ЕМ69	HE8EZ48	HEBGF73
Gene No.	51	52	53	54	55	26

Last AA of ORF	95	94	26	10	64	21
Last AA First AA of of Sig Secreted Pep Portion (36	30	22		20	22
Last AA of Sig Pep	35	29	21		61	21
irst AA of of Sig		_	_	_	-	-
SEQ SEQ ≺	181	182	183	184	185	186
of AA For SEQ AA of DO Signal NO: 9	459	63	839	270	272	127
of Start odor	459	63	839	270	272	127
South Seq. Seq. Seq. Complements Seq. Seq. Seq. Seq. Seq. Seq. Seq. Seq.	1090	260	1581	711	935	484
5' NT of Clone Seq.	267		765	∞	=	113
Total NT Seq.	1193	560	1657	7111	935	504
SEQ NO:	<i>L</i> 9	89	69	70	71	72
Vector	Uni-ZAP XR	Uni-ZAP XR	pBluescript	Lambda ZAP II	Lambda ZAP II	Lambda ZAP II
ATCC Deposit Nr and Date	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97
cDNA Clone ID	HFEBF41	HFRBU14	HFVGZ79	нндсм76	HHGCO88	нндсР52
Gene No.	57	58	59	09	61	62

Last AA of ORF	131	89	44	64	22	169
Last AA First AA 1 of of Sig Secreted Pep Portion C	61	33	28	37	12	15
Last AA of Sig Pep	18	32	27	36	=	14
irst AA of Sig						-
SEQ NÖ:	187	188	189	061	228	192
S' NT Of AA F First SEQ Of AA of AA of B Start Signal NO: Codon Pep Y I	96	248	630	167	575	187
5' NT of Start Codon	96	248	630	167		187
3' NT of Clone Seq.	620	581	1786	008	1076	1888
Seq. Seq.	-	156	537	116	398	18
Total NT Seq.	620	581	1843	1441	1076	2776
SEQ XÖ: BÖ:	73	74	75	92	114	78
Vector	Lambda ZAP II	Lambda ZAP II	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	pCMVSport 3.0
ATCC Deposit Nr and Date	97958 03/13/97 209072	97958 03/13/97 209072	97958 03/13/97 209072	97958 97958 03/13/97 209072	97958 03/13/97 209072 05/22/97	97958 . 97958 . 03/13/97 209072 05/22/97
cDNA Clone ID	HHGDB72	HHGDI71	HHSDI45	HHSEB66	HJPAZ83	HLDBO49
Gene No.	63	64	65	99	- 67	89

Last AA of ORF	65	131	91	175	69	24	72
First AA of Secreted Portion	23	23	33	24	27	21	26
Last AA of Sig Pep	22	22	32	23	26	20	25
First AA of Sig Pep		_	_	_		-	_
SEQ NO:	193	229	194	195	196	197	861
5' NT of First AA of Signal Pep	534	534	40	238	286	28	4
5' NT of Start Codon	534	534	40	238	286	58	41
S' NT 3' NT of of Of of Clone Clone NT Seq. Seq.	1480	1487	1077	780	770	481	623
S' NT of Clone Seq.	401	401	33	8	101	_	_
I	1525	1487	1563	1020	770	481	644
SEQ NÖ BÖ	62	115	08	8	82	83	84
Vector	pCMVSport 3.0	pCMVSport 3.0	Uni-ZAP XR	Uni-Zap XR	Uni-ZAP XR	Uni-ZAP XR	pCMVSport 3.0
ATCC Deposit Nr and Date	97958 03/13/97 209072 05/22/97	209226 08/28/97	97958 03/13/97 209072 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97
cDNA Clone ID	HLDBQ19	HLDBQ19	HMSGT42			HNGJU84	HNTAC73
Gene No.	69	69	70	71	72	73	74

	288	17	623	09	648	28
First AA of Secreted Portion	13		31	33	31	22
	12		30	32	30	21
AA AA of Sig Pep	1	-	_			_
SEQ NÖ:	199	230	200	231	201	232
S' NT of AA For SEQ AA of D Signal NO: Pep Y I	86	545	99	477	251	212
S' NT of Start Codon	86		99	477	251	677
3' NT of Clone Seq.	1284	1283	1747	1747	2566	1098
S' NT 3' NT of of Clone Clone Seq. Seq.	435	428	290	288	1843	375
Total NT Seq.	1351	1350	2527	2527	2566	1098
SEQ X	85	911	98	117	87	118
Vector	Uni-ZAP XR					
ATCC Deposit Nr and Date	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/27/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 97957 03/13/97 209073 05/22/97
cDNA Clone ID	HOSE145	HOSEI45	HOSFD58	HOSFD58	HSAUM95	HSAUM95
Gene No.	75	75	76	92	77	77

Last AA of ORF	54	265	17	314	206	194
First AA of Secreted Portion	33	12		20	21	70
Last AA of Sig Pep	32			61	20	69
First AA of Sig Pep		_	_	_	_	_
¥ŠeŠe ¥Še	202	203	233	204	205	206
5' NT of First AA of Signal Pep	83	188	315	92	414	157
5' NT of AA For S' NT Signal NO: Start Start Signal NO: Start Signal NO: Start Start Signal NO: Start Start Start Signal NO: Start Sta	83	188	315	92	414	157
5' NT 3' NT of of Clone Clone Seq. Seq.	540	1165	1166	2449	2058	1411
5' NT 3' NT of of of Clone Clone NT Seq. Seq.	<u> </u>	152	152	1149	476	345
	540	1863	1679	2478	2058	1411
SEQ NÖ: NÖ:	88	68	119	06	16	92
Vector	Uni-ZAP XR	pBluescript				
ATCC Deposit Nr and Date	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97
cDNA Clone ID	HSAUR67	HSKDI81	HSKDI81	HSKDW91	HTLEX50	HSKHL65
Gene No.	78	79	62	80	81	82

, , ,	71	329	95	57	391	25
First AA of Secreted Portion	38	31	20	21	2	22
Last AA of Sig Pep	37	30	61	20	_	21
st f f g	_		1		_	
AA SEQ ID NO: Y	235	207	236	208	209	210
5' NT AA Fir Of AA Fir AA of ID O Signal NO: Si Pep Y Pe	526	397	228	445	523	117
of of Star Sode	526	397	228	445	523	117
3' NT of Clone Seq.	1411	2184	2063	809	2394	672
S' NT 3' NT of of State Clone Clone NT Seq. Seq.	345	147	138	524	481	_
Total NT Seq.	1411	2187	2256	757	2394	672
SEQ NO:	121	93	122	94	95	96
Vector	pBluescript	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	ZAP Express	Uni-ZAP XR
ATCC Deposit Nr and Date	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/27/97	97957 03/13/97 209073 05/27/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/2/97	97957 03/13/97 209073 05/22/97
cDNA Clone ID	HSKHL65	HHFGA11	HHFGA11	HWTBL40	HBXFG80	HCACY32
Gene No.	82	83	83	84	82	98

Last AA of ORF	37
5' NT 3' NTof of o	21
Last AA of Sig Pep	20
First AA of Sig Pep	211 1 20
AA SEQ ID NO: Y	211
5' NT of First AA of Signal Pep	207
5' NT of Start Codon	207
3' NT of Clone Seq.	1419
5' NT of Clone Seq.	-
Total NT Seq.	1419
SEQ NO: NO:	97
Vector	97957 Uni-ZAP XR 97 1419 1 3/13/97 209073 5/22/97
ATCC Deposit Nr and Date	97957 03/13/97 209073 05/22/97
1	HCED021
Gene No.	87

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Table 1 summarizes the information corresponding to each "Gene No." described above. The nucleotide sequence identified as "NT SEQ ID NO:X" was assembled from partially homologous ("overlapping") sequences obtained from the "cDNA clone ID" identified in Table 1 and, in some cases, from additional related DNA clones. The overlapping sequences were assembled into a single contiguous sequence of high redundancy (usually three to five overlapping sequences at each nucleotide position), resulting in a final sequence identified as SEQ ID NO:X.

The cDNA Clone ID was deposited on the date and given the corresponding deposit number listed in "ATCC Deposit No:Z and Date." Some of the deposits contain multiple different clones corresponding to the same gene. "Vector" refers to the type of vector contained in the cDNA Clone ID.

"Total NT Seq." refers to the total number of nucleotides in the contig identified by "Gene No." The deposited clone may contain all or most of these sequences, reflected by the nucleotide position indicated as "5' NT of Clone Seq." and the "3' NT of Clone Seq." of SEQ ID NO:X. The nucleotide position of SEQ ID NO:X of the putative start codon (methionine) is identified as "5' NT of Start Codon." Similarly, the nucleotide position of SEQ ID NO:X of the predicted signal sequence is identified as "5' NT of First AA of Signal Pep."

The translated amino acid sequence, beginning with the methionine, is identified as "AA SEQ ID NO:Y," although other reading frames can also be easily translated using known molecular biology techniques. The polypeptides produced by these alternative open reading frames are specifically contemplated by the present invention.

The first and last amino acid position of SEQ ID NO:Y of the predicted signal peptide is identified as "First AA of Sig Pep" and "Last AA of Sig Pep." The predicted first amino acid position of SEQ ID NO:Y of the secreted portion is identified as "Predicted First AA of Secreted Portion." Finally, the amino acid position of SEQ ID NO:Y of the last amino acid in the open reading frame is identified as "Last AA of ORF."

SEQ ID NO:X and the translated SEQ ID NO:Y are sufficiently accurate and otherwise suitable for a variety of uses well known in the art and described further below. For instance, SEQ ID NO:X is useful for designing nucleic acid hybridization probes that will detect nucleic acid sequences contained in SEQ ID NO:X or the cDNA contained in the deposited clone. These probes will also hybridize to nucleic acid molecules in biological samples, thereby enabling a variety of forensic and diagnostic methods of the invention. Similarly, polypeptides identified from SEQ ID NO:Y may be used to generate antibodies which bind specifically to the secreted proteins encoded by the cDNA clones identified in Table 1.

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Nevertheless, DNA sequences generated by sequencing reactions can contain sequencing errors. The errors exist as misidentified nucleotides, or as insertions or deletions of nucleotides in the generated DNA sequence. The erroneously inserted or deleted nucleotides cause frame shifts in the reading frames of the predicted amino acid sequence. In these cases, the predicted amino acid sequence diverges from the actual amino acid sequence, even though the generated DNA sequence may be greater than 99.9% identical to the actual DNA sequence (for example, one base insertion or deletion in an open reading frame of over 1000 bases).

Accordingly, for those applications requiring precision in the nucleotide sequence or the amino acid sequence, the present invention provides not only the generated nucleotide sequence identified as SEQ ID NO:X and the predicted translated amino acid sequence identified as SEQ ID NO:Y, but also a sample of plasmid DNA containing a human cDNA of the invention deposited with the ATCC, as set forth in Table 1. The nucleotide sequence of each deposited clone can readily be determined by sequencing the deposited clone in accordance with known methods. The predicted amino acid sequence can then be verified from such deposits. Moreover, the amino acid sequence of the protein encoded by a particular clone can also be directly determined by peptide sequencing or by expressing the protein in a suitable host cell containing the deposited human cDNA, collecting the protein, and determining its sequence.

The present invention also relates to the genes corresponding to SEQ ID NO:X, SEQ ID NO:Y, or the deposited clone. The corresponding gene can be isolated in accordance with known methods using the sequence information disclosed herein. Such methods include preparing probes or primers from the disclosed sequence and identifying or amplifying the corresponding gene from appropriate sources of genomic material.

Also provided in the present invention are species homologs. Species homologs may be isolated and identified by making suitable probes or primers from the sequences provided herein and screening a suitable nucleic acid source for the desired homologue.

The polypeptides of the invention can be prepared in any suitable manner. Such polypeptides include isolated naturally occurring polypeptides, recombinantly produced polypeptides, synthetically produced polypeptides, or polypeptides produced by a combination of these methods. Means for preparing such polypeptides are well understood in the art.

The polypeptides may be in the form of the secreted protein, including the mature form, or may be a part of a larger protein, such as a fusion protein (see below).

It is often advantageous to include an additional amino acid sequence which contains secretory or leader sequences, pro-sequences, sequences which aid in purification, such as multiple histidine residues, or an additional sequence for stability during recombinant production.

The polypeptides of the present invention are preferably provided in an isolated form, and preferably are substantially purified. A recombinantly produced version of a polypeptide, including the secreted polypeptide, can be substantially purified by the one-step method described in Smith and Johnson, Gene 67:31-40 (1988). Polypeptides of the invention also can be purified from natural or recombinant sources using antibodies of the invention raised against the secreted protein in methods which are well known in the art.

Signal Sequences

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Methods for predicting whether a protein has a signal sequence, as well as the cleavage point for that sequence, are available. For instance, the method of McGeoch, Virus Res. 3:271-286 (1985), uses the information from a short N-terminal charged region and a subsequent uncharged region of the complete (uncleaved) protein. The method of von Heinje, Nucleic Acids Res. 14:4683-4690 (1986) uses the information from the residues surrounding the cleavage site, typically residues -13 to +2, where +1 indicates the amino terminus of the secreted protein. The accuracy of predicting the cleavage points of known mammalian secretory proteins for each of these methods is in the range of 75-80%. (von Heinje, supra.) However, the two methods do not always produce the same predicted cleavage point(s) for a given protein.

In the present case, the deduced amino acid sequence of the secreted polypeptide was analyzed by a computer program called SignalP (Henrik Nielsen et al., Protein Engineering 10:1-6 (1997)), which predicts the cellular location of a protein based on the amino acid sequence. As part of this computational prediction of localization, the methods of McGeoch and von Heinje are incorporated. The analysis of the amino acid sequences of the secreted proteins described herein by this program provided the results shown in Table 1.

As one of ordinary skill would appreciate, however, cleavage sites sometimes vary from organism to organism and cannot be predicted with absolute certainty. Accordingly, the present invention provides secreted polypeptides having a sequence shown in SEQ ID NO:Y which have an N-terminus beginning within 5 residues (i.e., + or - 5 residues) of the predicted cleavage point. Similarly, it is also recognized that in some cases, cleavage of the signal sequence from a secreted protein is not entirely

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uniform, resulting in more than one secreted species. These polypeptides, and the polynucleotides encoding such polypeptides, are contemplated by the present invention.

Moreover, the signal sequence identified by the above analysis may not necessarily predict the naturally occurring signal sequence. For example, the naturally occurring signal sequence may be further upstream from the predicted signal sequence. However, it is likely that the predicted signal sequence will be capable of directing the secreted protein to the ER. These polypeptides, and the polynucleotides encoding such polypeptides, are contemplated by the present invention.

10 Polynucleotide and Polypeptide Variants

"Variant" refers to a polynucleotide or polypeptide differing from the polynucleotide or polypeptide of the present invention, but retaining essential properties thereof. Generally, variants are overall closely similar, and, in many regions, identical to the polynucleotide or polypeptide of the present invention.

By a polynucleotide having a nucleotide sequence at least, for example, 95% "identical" to a reference nucleotide sequence of the present invention, it is intended that the nucleotide sequence of the polynucleotide is identical to the reference sequence except that the polynucleotide sequence may include up to five point mutations per each 100 nucleotides of the reference nucleotide sequence encoding the polypeptide. In other words, to obtain a polynucleotide having a nucleotide sequence at least 95% identical to a reference nucleotide sequence, up to 5% of the nucleotides in the reference sequence may be deleted or substituted with another nucleotide, or a number of nucleotides up to 5% of the total nucleotides in the reference sequence may be inserted into the reference sequence. The query sequence may be an entire sequence shown inTable 1, the ORF (open reading frame), or any fragement specified as described herein.

As a practical matter, whether any particular nucleic acid molecule or polypeptide is at least 90%, 95%, 96%, 97%, 98% or 99% identical to a nucleotide sequence of the presence invention can be determined conventionally using known computer programs. A preferred method for determing the best overall match between a query sequence (a sequence of the present invention) and a subject sequence, also referred to as a global sequence alignment, can be determined using the FASTDB computer program based on the algorithm of Brutlag et al. (Comp. App. Biosci. (1990) 6:237-245). In a sequence alignment the query and subject sequences are both DNA sequences. An RNA sequence can be compared by converting U's to T's. The result of said global sequence alignment is in percent identity. Preferred parameters used in a FASTDB alignment of DNA sequences to calculate percent identity are:

Matrix=Unitary, k-tuple=4, Mismatch Penalty=1, Joining Penalty=30, Randomization

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Group Length=0, Cutoff Score=1, Gap Penalty=5, Gap Size Penalty 0.05, Window Size=500 or the length of the subject nucleotide sequence, whichever is shorter.

If the subject sequence is shorter than the query sequence because of 5' or 3' deletions, not because of internal deletions, a manual correction must be made to the results. This is because the FASTDB program does not account for 5' and 3' truncations of the subject sequence when calculating percent identity. For subject sequences truncated at the 5' or 3' ends, relative to the the query sequence, the percent identity is corrected by calculating the number of bases of the query sequence that are 5' and 3' of the subject sequence, which are not matched/aligned, as a percent of the total bases of the query sequence. Whether a nucleotide is matched/aligned is determined by results of the FASTDB sequence alignment. This percentage is then subtracted from the percent identity, calculated by the above FASTDB program using the specified parameters, to arrive at a final percent identity score. This corrected score is what is used for the purposes of the present invention. Only bases outside the 5' and 3' bases of the subject sequence, as displayed by the FASTDB alignment, which are not matched/aligned with the query sequence, are calculated for the purposes of manually adjusting the percent identity score.

For example, a 90 base subject sequence is aligned to a 100 base query sequence to determine percent identity. The deletions occur at the 5' end of the subject sequence and therefore, the FASTDB alignment does not show a matched/alignement of the first 10 bases at 5' end. The 10 unpaired bases represent 10% of the sequence (number of bases at the 5' and 3' ends not matched/total number of bases in the query sequence) so 10% is subtracted from the percent identity score calculated by the FASTDB program. If the remaining 90 bases were perfectly matched the final percent identity would be 90%. In another example, a 90 base subject sequence is compared with a 100 base query sequence. This time the deletions are internal deletions so that there are no bases on the 5' or 3' of the subject sequence which are not matched/aligned with the query. In this case the percent identity calculated by FASTDB is not manually corrected. Once again, only bases 5' and 3' of the subject sequence which are not matched/aligned with the query sequence are manually corrected for. No other manual corrections are to made for the purposes of the present invention.

By a polypeptide having an amino acid sequence at least, for example, 95% "identical" to a query amino acid sequence of the present invention, it is intended that the amino acid sequence of the subject polypeptide is identical to the query sequence except that the subject polypeptide sequence may include up to five amino acid alterations per each 100 amino acids of the query amino acid sequence. In other words, to obtain a polypeptide having an amino acid sequence at least 95% identical to a query

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amino acid sequence, up to 5% of the amino acid residues in the subject sequence may be inserted, deleted, (indels) or substituted with another amino acid. These alterations of the reference sequence may occur at the amino or carboxy terminal positions of the reference amino acid sequence or anywhere between those terminal positions, interspersed either individually among residues in the reference sequence or in one or more contiguous groups within the reference sequence.

As a practical matter, whether any particular polypeptide is at least 90%, 95%, 96%, 97%, 98% or 99% identical to, for instance, the amino acid sequences shown in Table 1 or to the amino acid sequence encoded by deposited DNA clone can be determined conventionally using known computer programs. A preferred method for determing the best overall match between a query sequence (a sequence of the present invention) and a subject sequence, also referred to as a global sequence alignment, can be determined using the FASTDB computer program based on the algorithm of Brutlag et al. (Comp. App. Biosci. (1990) 6:237-245). In a sequence alignment the query and subject sequences are either both nucleotide sequences or both amino acid sequences. The result of said global sequence alignment is in percent identity. Preferred parameters used in a FASTDB amino acid alignment are: Matrix=PAM 0, k-tuple=2, Mismatch Penalty=1, Joining Penalty=20, Randomization Group Length=0, Cutoff Score=1, Window Size=sequence length, Gap Penalty=5, Gap Size Penalty=0.05, Window Size=500 or the length of the subject amino acid sequence, whichever is shorter.

If the subject sequence is shorter than the query sequence due to N- or Cterminal deletions, not because of internal deletions, a manual correction must be made to the results. This is because the FASTDB program does not account for N- and Cterminal truncations of the subject sequence when calculating global percent identity. For subject sequences truncated at the N- and C-termini, relative to the the query sequence, the percent identity is corrected by calculating the number of residues of the query sequence that are N- and C-terminal of the subject sequence, which are not matched/aligned with a corresponding subject residue, as a percent of the total bases of the query sequence. Whether a residue is matched/aligned is determined by results of the FASTDB sequence alignment. This percentage is then subtracted from the percent identity, calculated by the above FASTDB program using the specified parameters, to arrive at a final percent identity score. This final percent identity score is what is used for the purposes of the present invention. Only residues to the N- and C-termini of the subject sequence, which are not matched/aligned with the query sequence, are considered for the purposes of manually adjusting the percent identity score. That is, only query residue positions outside the farthest N- and C-terminal residues of the subject sequence.

For example, a 90 amino acid residue subject sequence is aligned with a 100 residue query sequence to determine percent identity. The deletion occurs at the Nterminus of the subject sequence and therefore, the FASTDB alignment does not show a matching/alignment of the first 10 residues at the N-terminus. The 10 unpaired 5 residues represent 10% of the sequence (number of residues at the N- and C- termini not matched/total number of residues in the query sequence) so 10% is subtracted from the percent identity score calculated by the FASTDB program. If the remaining 90 residues were perfectly matched the final percent identity would be 90%. In another example, a 90 residue subject sequence is compared with a 100 residue query sequence. 10 This time the deletions are internal deletions so there are no residues at the N- or Ctermini of the subject sequence which are not matched/aligned with the query. In this case the percent identity calculated by FASTDB is not manually corrected. Once again, only residue positions outside the N- and C-terminal ends of the subject sequence, as displayed in the FASTDB alignment, which are not matched/aligned with the query sequnce are manually corrected for. No other manual corrections are to made for the 15 purposes of the present invention.

The variants may contain alterations in the coding regions, non-coding regions, or both. Especially preferred are polynucleotide variants containing alterations which produce silent substitutions, additions, or deletions, but do not alter the properties or activities of the encoded polypeptide. Nucleotide variants produced by silent substitutions due to the degeneracy of the genetic code are preferred. Moreover, variants in which 5-10, 1-5, or 1-2 amino acids are substituted, deleted, or added in any combination are also preferred. Polynucleotide variants can be produced for a variety of reasons, e.g., to optimize codon expression for a particular host (change codons in the human mRNA to those preferred by a bacterial host such as E. coli).

Naturally occurring variants are called "allelic variants," and refer to one of several alternate forms of a gene occupying a given locus on a chromosome of an organism. (Genes II, Lewin, B., ed., John Wiley & Sons, New York (1985).) These allelic variants can vary at either the polynucleotide and/or polypeptide level. Alternatively, non-naturally occurring variants may be produced by mutagenesis techniques or by direct synthesis.

Using known methods of protein engineering and recombinant DNA technology, variants may be generated to improve or alter the characteristics of the polypeptides of the present invention. For instance, one or more amino acids can be deleted from the N-terminus or C-terminus of the secreted protein without substantial loss of biological function. The authors of Ron et al., J. Biol. Chem. 268: 2984-2988 (1993), reported variant KGF proteins having heparin binding activity even after

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deleting 3, 8, or 27 amino-terminal amino acid residues. Similarly, Interferon gamma exhibited up to ten times higher activity after deleting 8-10 amino acid residues from the carboxy terminus of this protein. (Dobeli et al., J. Biotechnology 7:199-216 (1988).)

Moreover, ample evidence demonstrates that variants often retain a biological activity similar to that of the naturally occurring protein. For example, Gayle and coworkers (J. Biol. Chem 268:22105-22111 (1993)) conducted extensive mutational analysis of human cytokine IL-1a. They used random mutagenesis to generate over 3,500 individual IL-1a mutants that averaged 2.5 amino acid changes per variant over the entire length of the molecule. Multiple mutations were examined at every possible amino acid position. The investigators found that "[m]ost of the molecule could be altered with little effect on either [binding or biological activity]." (See, Abstract.) In fact, only 23 unique amino acid sequences, out of more than 3,500 nucleotide sequences examined, produced a protein that significantly differed in activity from wild-type.

Furthermore, even if deleting one or more amino acids from the N-terminus or C-terminus of a polypeptide results in modification or loss of one or more biological functions, other biological activities may still be retained. For example, the ability of a deletion variant to induce and/or to bind antibodies which recognize the secreted form will likely be retained when less than the majority of the residues of the secreted form are removed from the N-terminus or C-terminus. Whether a particular polypeptide lacking N- or C-terminal residues of a protein retains such immunogenic activities can readily be determined by routine methods described herein and otherwise known in the art.

Thus, the invention further includes polypeptide variants which show substantial biological activity. Such variants include deletions, insertions, inversions, repeats, and substitutions selected according to general rules known in the art so as have little effect on activity. For example, guidance concerning how to make phenotypically silent amino acid substitutions is provided in Bowie, J. U. et al., Science 247:1306-1310 (1990), wherein the authors indicate that there are two main strategies for studying the tolerance of an amino acid sequence to change.

The first strategy exploits the tolerance of amino acid substitutions by natural selection during the process of evolution. By comparing amino acid sequences in different species, conserved amino acids can be identified. These conserved amino acids are likely important for protein function. In contrast, the amino acid positions where substitutions have been tolerated by natural selection indicates that these positions are not critical for protein function. Thus, positions tolerating amino acid substitution could be modified while still maintaining biological activity of the protein.

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The second strategy uses genetic engineering to introduce amino acid changes at specific positions of a cloned gene to identify regions critical for protein function. For example, site directed mutagenesis or alanine-scanning mutagenesis (introduction of single alanine mutations at every residue in the molecule) can be used. (Cunningham and Wells, Science 244:1081-1085 (1989).) The resulting mutant molecules can then be tested for biological activity.

As the authors state, these two strategies have revealed that proteins are surprisingly tolerant of amino acid substitutions. The authors further indicate which amino acid changes are likely to be permissive at certain amino acid positions in the protein. For example, most buried (within the tertiary structure of the protein) amino acid residues require nonpolar side chains, whereas few features of surface side chains are generally conserved. Moreover, tolerated conservative amino acid substitutions involve replacement of the aliphatic or hydrophobic amino acids Ala, Val, Leu and Ile; replacement of the hydroxyl residues Ser and Thr; replacement of the acidic residues Asp and Glu; replacement of the amide residues Asn and Gln, replacement of the basic residues Lys, Arg, and His; replacement of the aromatic residues Phe, Tyr, and Trp, and replacement of the small-sized amino acids Ala, Ser, Thr, Met, and Gly.

Besides conservative amino acid substitution, variants of the present invention include (i) substitutions with one or more of the non-conserved amino acid residues, where the substituted amino acid residues may or may not be one encoded by the genetic code, or (ii) substitution with one or more of amino acid residues having a substituent group, or (iii) fusion of the mature polypeptide with another compound, such as a compound to increase the stability and/or solubility of the polypeptide (for example, polyethylene glycol), or (iv) fusion of the polypeptide with additional amino acids, such as an IgG Fc fusion region peptide, or leader or secretory sequence, or a sequence facilitating purification. Such variant polypeptides are deemed to be within the scope of those skilled in the art from the teachings herein.

For example, polypeptide variants containing amino acid substitutions of charged amino acids with other charged or neutral amino acids may produce proteins with improved characteristics, such as less aggregation. Aggregation of pharmaceutical formulations both reduces activity and increases clearance due to the aggregate's immunogenic activity. (Pinckard et al., Clin. Exp. Immunol. 2:331-340 (1967); Robbins et al., Diabetes 36: 838-845 (1987); Cleland et al., Crit. Rev. Therapeutic Drug Carrier Systems 10:307-377 (1993).)

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Polynucleotide and Polypeptide Fragments

In the present invention, a "polynucleotide fragment" refers to a short polynucleotide having a nucleic acid sequence contained in the deposited clone or shown in SEQ ID NO:X. The short nucleotide fragments are preferably at least about 15 nt, and more preferably at least about 20 nt, still more preferably at least about 30 nt, and even more preferably, at least about 40 nt in length. A fragment "at least 20 nt in length," for example, is intended to include 20 or more contiguous bases from the cDNA sequence contained in the deposited clone or the nucleotide sequence shown in SEQ ID NO:X. These nucleotide fragments are useful as diagnostic probes and primers as discussed herein. Of course, larger fragments (e.g., 50, 150, 500, 600, 2000 nucleotides) are preferred.

Moreover, representative examples of polynucleotide fragments of the invention, include, for example, fragments having a sequence from about nucleotide number 1-50, 51-100, 101-150, 151-200, 201-250, 251-300, 301-350, 351-400, 401-450, 451-500, 501-550, 551-600, 651-700, or 701 to the end of SEQ ID NO:X or the cDNA contained in the deposited clone. In this context "about" includes the particularly recited ranges, larger or smaller by several (5, 4, 3, 2, or 1) nucleotides, at either terminus or at both termini. Preferably, these fragments encode a polypeptide which has biological activity.

In the present invention, a "polypeptide fragment" refers to a short amino acid sequence contained in SEQ ID NO:Y or encoded by the cDNA contained in the deposited clone. Protein fragments may be "free-standing," or comprised within a larger polypeptide of which the fragment forms a part or region, most preferably as a single continuous region. Representative examples of polypeptide fragments of the invention, include, for example, fragments from about amino acid number 1-20, 21-40, 41-60, 61-80, 81-100, 102-120, 121-140, 141-160, or 161 to the end of the coding region. Moreover, polypeptide fragments can be about 20, 30, 40, 50, 60, 70, 80, 90, 100, 110, 120, 130, 140, or 150 amino acids in length. In this context "about" includes the particularly recited ranges, larger or smaller by several (5, 4, 3, 2, or 1) amino acids, at either extreme or at both extremes.

Preferred polypeptide fragments include the secreted protein as well as the mature form. Further preferred polypeptide fragments include the secreted protein or the mature form having a continuous series of deleted residues from the amino or the carboxy terminus, or both. For example, any number of amino acids, ranging from 1-60, can be deleted from the amino terminus of either the secreted polypeptide or the mature form. Similarly, any number of amino acids, ranging from 1-30, can be deleted from the carboxy terminus of the secreted protein or mature form. Furthermore, any

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combination of the above amino and carboxy terminus deletions are preferred. Similarly, polynucleotide fragments encoding these polypeptide fragments are also preferred.

Also preferred are polypeptide and polynucleotide fragments characterized by structural or functional domains, such as fragments that comprise alpha-helix and alpha-helix forming regions, beta-sheet and beta-sheet-forming regions, turn and turn-forming regions, coil and coil-forming regions, hydrophilic regions, hydrophobic regions, alpha amphipathic regions, beta amphipathic regions, flexible regions, surface-forming regions, substrate binding region, and high antigenic index regions. Polypeptide fragments of SEQ ID NO:Y falling within conserved domains are specifically contemplated by the present invention. Moreover, polynucleotide fragments encoding these domains are also contemplated.

Other preferred fragments are biologically active fragments. Biologically active fragments are those exhibiting activity similar, but not necessarily identical, to an activity of the polypeptide of the present invention. The biological activity of the fragments may include an improved desired activity, or a decreased undesirable activity.

Epitopes & Antibodies

In the present invention, "epitopes" refer to polypeptide fragments having antigenic or immunogenic activity in an animal, especially in a human. A preferred embodiment of the present invention relates to a polypeptide fragment comprising an epitope, as well as the polynucleotide encoding this fragment. A region of a protein molecule to which an antibody can bind is defined as an "antigenic epitope." In contrast, an "immunogenic epitope" is defined as a part of a protein that elicits an antibody response. (See, for instance, Geysen et al., Proc. Natl. Acad. Sci. USA 81:3998- 4002 (1983).)

Fragments which function as epitopes may be produced by any conventional means. (See, e.g., Houghten, R. A., Proc. Natl. Acad. Sci. USA 82:5131-5135 (1985) further described in U.S. Patent No. 4,631,211.)

In the present invention, antigenic epitopes preferably contain a sequence of at least seven, more preferably at least nine, and most preferably between about 15 to about 30 amino acids. Antigenic epitopes are useful to raise antibodies, including monoclonal antibodies, that specifically bind the epitope. (See, for instance, Wilson et al., Cell 37:767-778 (1984); Sutcliffe, J. G. et al., Science 219:660-666 (1983).)

Similarly, immunogenic epitopes can be used to induce antibodies according to methods well known in the art. (See, for instance, Sutcliffe et al., supra; Wilson et al., supra; Chow, M. et al., Proc. Natl. Acad. Sci. USA 82:910-914; and Bittle, F. J. et

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al., J. Gen. Virol. 66:2347-2354 (1985).) A preferred immunogenic epitope includes the secreted protein. The immunogenic epitopes may be presented together with a carrier protein, such as an albumin, to an animal system (such as rabbit or mouse) or, if it is long enough (at least about 25 amino acids), without a carrier. However, immunogenic epitopes comprising as few as 8 to 10 amino acids have been shown to be sufficient to raise antibodies capable of binding to, at the very least, linear epitopes in a denatured polypeptide (e.g., in Western blotting.)

As used herein, the term "antibody" (Ab) or "monoclonal antibody" (Mab) is meant to include intact molecules as well as antibody fragments (such as, for example, Fab and F(ab')2 fragments) which are capable of specifically binding to protein. Fab and F(ab')2 fragments lack the Fc fragment of intact antibody, clear more rapidly from the circulation, and may have less non-specific tissue binding than an intact antibody. (Wahl et al., J. Nucl. Med. 24:316-325 (1983).) Thus, these fragments are preferred, as well as the products of a FAB or other immunoglobulin expression library. Moreover, antibodies of the present invention include chimeric, single chain, and humanized antibodies.

Fusion Proteins

Any polypeptide of the present invention can be used to generate fusion proteins. For example, the polypeptide of the present invention, when fused to a second protein, can be used as an antigenic tag. Antibodies raised against the polypeptide of the present invention can be used to indirectly detect the second protein by binding to the polypeptide. Moreover, because secreted proteins target cellular locations based on trafficking signals, the polypeptides of the present invention can be used as targeting molecules once fused to other proteins.

Examples of domains that can be fused to polypeptides of the present invention include not only heterologous signal sequences, but also other heterologous functional regions. The fusion does not necessarily need to be direct, but may occur through linker sequences.

Moreover, fusion proteins may also be engineered to improve characteristics of the polypeptide of the present invention. For instance, a region of additional amino acids, particularly charged amino acids, may be added to the N-terminus of the polypeptide to improve stability and persistence during purification from the host cell or subsequent handling and storage. Also, peptide moieties may be added to the polypeptide to facilitate purification. Such regions may be removed prior to final preparation of the polypeptide. The addition of peptide moieties to facilitate handling of polypeptides are familiar and routine techniques in the art.

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Moreover, polypeptides of the present invention, including fragments, and specifically epitopes, can be combined with parts of the constant domain of immunoglobulins (IgG), resulting in chimeric polypeptides. These fusion proteins facilitate purification and show an increased half-life in vivo. One reported example describes chimeric proteins consisting of the first two domains of the human CD4-polypeptide and various domains of the constant regions of the heavy or light chains of mammalian immunoglobulins. (EP A 394,827; Traunecker et al., Nature 331:84-86 (1988).) Fusion proteins having disulfide-linked dimeric structures (due to the IgG) can also be more efficient in binding and neutralizing other molecules, than the monomeric secreted protein or protein fragment alone. (Fountoulakis et al., J. Biochem. 270:3958-3964 (1995).)

Similarly, EP-A-O 464 533 (Canadian counterpart 2045869) discloses fusion proteins comprising various portions of constant region of immunoglobulin molecules together with another human protein or part thereof. In many cases, the Fc part in a fusion protein is beneficial in therapy and diagnosis, and thus can result in, for example, improved pharmacokinetic properties. (EP-A 0232 262.) Alternatively, deleting the Fc part after the fusion protein has been expressed, detected, and purified, would be desired. For example, the Fc portion may hinder therapy and diagnosis if the fusion protein is used as an antigen for immunizations. In drug discovery, for example, human proteins, such as hIL-5, have been fused with Fc portions for the purpose of high-throughput screening assays to identify antagonists of hIL-5. (See, D. Bennett et al., J. Molecular Recognition 8:52-58 (1995); K. Johanson et al., J. Biol. Chem. 270:9459-9471 (1995).)

Moreover, the polypeptides of the present invention can be fused to marker sequences, such as a peptide which facilitates purification of the fused polypeptide. In preferred embodiments, the marker amino acid sequence is a hexa-histidine peptide, such as the tag provided in a pQE vector (QIAGEN, Inc., 9259 Eton Avenue, Chatsworth, CA, 91311), among others, many of which are commercially available. As described in Gentz et al., Proc. Natl. Acad. Sci. USA 86:821-824 (1989), for instance, hexa-histidine provides for convenient purification of the fusion protein. Another peptide tag useful for purification, the "HA" tag, corresponds to an epitope derived from the influenza hemagglutinin protein. (Wilson et al., Cell 37:767 (1984).)

Thus, any of these above fusions can be engineered using the polynucleotides or the polypeptides of the present invention.

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Vectors, Host Cells, and Protein Production

The present invention also relates to vectors containing the polynucleotide of the present invention, host cells, and the production of polypeptides by recombinant techniques. The vector may be, for example, a phage, plasmid, viral, or retroviral vector. Retroviral vectors may be replication competent or replication defective. In the latter case, viral propagation generally will occur only in complementing host cells.

The polynucleotides may be joined to a vector containing a selectable marker for propagation in a host. Generally, a plasmid vector is introduced in a precipitate, such as a calcium phosphate precipitate, or in a complex with a charged lipid. If the vector is a virus, it may be packaged in vitro using an appropriate packaging cell line and then transduced into host cells.

The polynucleotide insert should be operatively linked to an appropriate promoter, such as the phage lambda PL promoter, the E. coli lac, trp, phoA and tac promoters, the SV40 early and late promoters and promoters of retroviral LTRs, to name a few. Other suitable promoters will be known to the skilled artisan. The expression constructs will further contain sites for transcription initiation, termination, and, in the transcribed region, a ribosome binding site for translation. The coding portion of the transcripts expressed by the constructs will preferably include a translation initiating codon at the beginning and a termination codon (UAA, UGA or UAG) appropriately positioned at the end of the polypeptide to be translated.

As indicated, the expression vectors will preferably include at least one selectable marker. Such markers include dihydrofolate reductase, G418 or neomycin resistance for eukaryotic cell culture and tetracycline, kanamycin or ampicillin resistance genes for culturing in E. coli and other bacteria. Representative examples of appropriate hosts include, but are not limited to, bacterial cells, such as E. coli, Streptomyces and Salmonella typhimurium cells; fungal cells, such as yeast cells; insect cells such as Drosophila S2 and Spodoptera Sf9 cells; animal cells such as CHO, COS, 293, and Bowes melanoma cells; and plant cells. Appropriate culture mediums and conditions for the above-described host cells are known in the art.

Among vectors preferred for use in bacteria include pQE70, pQE60 and pQE-9, available from QIAGEN, Inc.; pBluescript vectors, Phagescript vectors, pNH8A, pNH16a, pNH18A, pNH46A, available from Stratagene Cloning Systems, Inc.; and ptrc99a, pKK223-3, pKK233-3, pDR540, pRIT5 available from Pharmacia Biotech, Inc. Among preferred eukaryotic vectors are pWLNEO, pSV2CAT, pOG44, pXT1 and pSG available from Stratagene; and pSVK3, pBPV, pMSG and pSVL available from Pharmacia. Other suitable vectors will be readily apparent to the skilled artisan.

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Introduction of the construct into the host cell can be effected by calcium phosphate transfection, DEAE-dextran mediated transfection, cationic lipid-mediated transfection, electroporation, transduction, infection, or other methods. Such methods are described in many standard laboratory manuals, such as Davis et al., Basic Methods In Molecular Biology (1986). It is specifically contemplated that the polypeptides of the present invention may in fact be expressed by a host cell lacking a recombinant vector.

A polypeptide of this invention can be recovered and purified from recombinant cell cultures by well-known methods including ammonium sulfate or ethanol precipitation, acid extraction, anion or cation exchange chromatography, phosphocellulose chromatography, hydrophobic interaction chromatography, affinity chromatography, hydroxylapatite chromatography and lectin chromatography. Most preferably, high performance liquid chromatography ("HPLC") is employed for purification.

Polypeptides of the present invention, and preferably the secreted form, can also be recovered from: products purified from natural sources, including bodily fluids, 15 tissues and cells, whether directly isolated or cultured; products of chemical synthetic procedures; and products produced by recombinant techniques from a prokaryotic or eukaryotic host, including, for example, bacterial, yeast, higher plant, insect, and mammalian cells. Depending upon the host employed in a recombinant production procedure, the polypeptides of the present invention may be glycosylated or may be 20 non-glycosylated. In addition, polypeptides of the invention may also include an initial modified methionine residue, in some cases as a result of host-mediated processes. Thus, it is well known in the art that the N-terminal methionine encoded by the translation initiation codon generally is removed with high efficiency from any protein after translation in all eukaryotic cells. While the N-terminal methionine on most 25 proteins also is efficiently removed in most prokaryotes, for some proteins, this prokaryotic removal process is inefficient, depending on the nature of the amino acid to which the N-terminal methionine is covalently linked.

30 Uses of the Polynucleotides

Each of the polynucleotides identified herein can be used in numerous ways as reagents. The following description should be considered exemplary and utilizes known techniques.

The polynucleotides of the present invention are useful for chromosome identification. There exists an ongoing need to identify new chromosome markers, since few chromosome marking reagents, based on actual sequence data (repeat

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polymorphisms), are presently available. Each polynucleotide of the present invention can be used as a chromosome marker.

Briefly, sequences can be mapped to chromosomes by preparing PCR primers (preferably 15-25 bp) from the sequences shown in SEQ ID NO:X. Primers can be selected using computer analysis so that primers do not span more than one predicted exon in the genomic DNA. These primers are then used for PCR screening of somatic cell hybrids containing individual human chromosomes. Only those hybrids containing the human gene corresponding to the SEQ ID NO:X will yield an amplified fragment.

Similarly, somatic hybrids provide a rapid method of PCR mapping the polynucleotides to particular chromosomes. Three or more clones can be assigned per day using a single thermal cycler. Moreover, sublocalization of the polynucleotides can be achieved with panels of specific chromosome fragments. Other gene mapping strategies that can be used include in situ hybridization, prescreening with labeled flow-sorted chromosomes, and preselection by hybridization to construct chromosome specific-cDNA libraries.

Precise chromosomal location of the polynucleotides can also be achieved using fluorescence in situ hybridization (FISH) of a metaphase chromosomal spread. This technique uses polynucleotides as short as 500 or 600 bases; however, polynucleotides 2,000-4,000 bp are preferred. For a review of this technique, see Verma et al., "Human Chromosomes: a Manual of Basic Techniques," Pergamon Press, New York (1988).

For chromosome mapping, the polynucleotides can be used individually (to mark a single chromosome or a single site on that chromosome) or in panels (for marking multiple sites and/or multiple chromosomes). Preferred polynucleotides correspond to the noncoding regions of the cDNAs because the coding sequences are more likely conserved within gene families, thus increasing the chance of cross hybridization during chromosomal mapping.

Once a polynucleotide has been mapped to a precise chromosomal location, the physical position of the polynucleotide can be used in linkage analysis. Linkage analysis establishes coinheritance between a chromosomal location and presentation of a particular disease. (Disease mapping data are found, for example, in V. McKusick, Mendelian Inheritance in Man (available on line through Johns Hopkins University Welch Medical Library).) Assuming 1 megabase mapping resolution and one gene per 20 kb, a cDNA precisely localized to a chromosomal region associated with the disease could be one of 50-500 potential causative genes.

Thus, once coinheritance is established, differences in the polynucleotide and the corresponding gene between affected and unaffected individuals can be examined.

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First, visible structural alterations in the chromosomes, such as deletions or translocations, are examined in chromosome spreads or by PCR. If no structural alterations exist, the presence of point mutations are ascertained. Mutations observed in some or all affected individuals, but not in normal individuals, indicates that the mutation may cause the disease. However, complete sequencing of the polypeptide and the corresponding gene from several normal individuals is required to distinguish the mutation from a polymorphism. If a new polymorphism is identified, this polymorphic polypeptide can be used for further linkage analysis.

Furthermore, increased or decreased expression of the gene in affected individuals as compared to unaffected individuals can be assessed using polynucleotides of the present invention. Any of these alterations (altered expression, chromosomal rearrangement, or mutation) can be used as a diagnostic or prognostic marker.

In addition to the foregoing, a polynucleotide can be used to control gene expression through triple helix formation or antisense DNA or RNA. Both methods rely on binding of the polynucleotide to DNA or RNA. For these techniques, preferred polynucleotides are usually 20 to 40 bases in length and complementary to either the region of the gene involved in transcription (triple helix - see Lee et al., Nucl. Acids Res. 6:3073 (1979); Cooney et al., Science 241:456 (1988); and Dervan et al., Science 251:1360 (1991)) or to the mRNA itself (antisense - Okano, J. Neurochem. 56:560 (1991); Oligodeoxy-nucleotides as Antisense Inhibitors of Gene Expression, CRC Press, Boca Raton, FL (1988).) Triple helix formation optimally results in a shut-off of RNA transcription from DNA, while antisense RNA hybridization blocks translation of an mRNA molecule into polypeptide. Both techniques are effective in model systems, and the information disclosed herein can be used to design antisense or triple helix polynucleotides in an effort to treat disease.

Polynucleotides of the present invention are also useful in gene therapy. One goal of gene therapy is to insert a normal gene into an organism having a defective gene, in an effort to correct the genetic defect. The polynucleotides disclosed in the present invention offer a means of targeting such genetic defects in a highly accurate manner. Another goal is to insert a new gene that was not present in the host genome, thereby producing a new trait in the host cell.

The polynucleotides are also useful for identifying individuals from minute biological samples. The United States military, for example, is considering the use of restriction fragment length polymorphism (RFLP) for identification of its personnel. In this technique, an individual's genomic DNA is digested with one or more restriction enzymes, and probed on a Southern blot to yield unique bands for identifying

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personnel. This method does not suffer from the current limitations of "Dog Tags" which can be lost, switched, or stolen, making positive identification difficult. The polynucleotides of the present invention can be used as additional DNA markers for RFLP.

The polynucleotides of the present invention can also be used as an alternative to RFLP, by determining the actual base-by-base DNA sequence of selected portions of an individual's genome. These sequences can be used to prepare PCR primers for amplifying and isolating such selected DNA, which can then be sequenced. Using this technique, individuals can be identified because each individual will have a unique set of DNA sequences. Once an unique ID database is established for an individual, positive identification of that individual, living or dead, can be made from extremely small tissue samples.

Forensic biology also benefits from using DNA-based identification techniques as disclosed herein. DNA sequences taken from very small biological samples such as tissues, e.g., hair or skin, or body fluids, e.g., blood, saliva, semen, etc., can be amplified using PCR. In one prior art technique, gene sequences amplified from polymorphic loci, such as DQa class II HLA gene, are used in forensic biology to identify individuals. (Erlich, H., PCR Technology, Freeman and Co. (1992).) Once these specific polymorphic loci are amplified, they are digested with one or more restriction enzymes, yielding an identifying set of bands on a Southern blot probed with DNA corresponding to the DQa class II HLA gene. Similarly, polynucleotides of the present invention can be used as polymorphic markers for forensic purposes.

There is also a need for reagents capable of identifying the source of a particular tissue. Such need arises, for example, in forensics when presented with tissue of unknown origin. Appropriate reagents can comprise, for example, DNA probes or primers specific to particular tissue prepared from the sequences of the present invention. Panels of such reagents can identify tissue by species and/or by organ type. In a similar fashion, these reagents can be used to screen tissue cultures for contamination.

In the very least, the polynucleotides of the present invention can be used as molecular weight markers on Southern gels, as diagnostic probes for the presence of a specific mRNA in a particular cell type, as a probe to "subtract-out" known sequences in the process of discovering novel polynucleotides, for selecting and making oligomers for attachment to a "gene chip" or other support, to raise anti-DNA antibodies using DNA immunization techniques, and as an antigen to elicit an immune response.

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Uses of the Polypeptides

Each of the polypeptides identified herein can be used in numerous ways. The following description should be considered exemplary and utilizes known techniques.

A polypeptide of the present invention can be used to assay protein levels in a biological sample using antibody-based techniques. For example, protein expression in tissues can be studied with classical immunohistological methods. (Jalkanen, M., et al., J. Cell. Biol. 101:976-985 (1985); Jalkanen, M., et al., J. Cell . Biol. 105:3087-3096 (1987).) Other antibody-based methods useful for detecting protein gene expression include immunoassays, such as the enzyme linked immunosorbent assay (ELISA) and the radioimmunoassay (RIA). Suitable antibody assay labels are known in the art and include enzyme labels, such as, glucose oxidase, and radioisotopes, such as iodine (125I, 121I), carbon (14C), sulfur (35S), tritium (3H), indium (112In), and technetium (99mTc), and fluorescent labels, such as fluorescein and rhodamine, and biotin.

In addition to assaying secreted protein levels in a biological sample, proteins can also be detected in vivo by imaging. Antibody labels or markers for in vivo imaging of protein include those detectable by X-radiography, NMR or ESR. For X-radiography, suitable labels include radioisotopes such as barium or cesium, which emit detectable radiation but are not overtly harmful to the subject. Suitable markers for NMR and ESR include those with a detectable characteristic spin, such as deuterium, which may be incorporated into the antibody by labeling of nutrients for the relevant hybridoma.

A protein-specific antibody or antibody fragment which has been labeled with an appropriate detectable imaging moiety, such as a radioisotope (for example, 131I, 112In, 99mTc), a radio-opaque substance, or a material detectable by nuclear magnetic resonance, is introduced (for example, parenterally, subcutaneously, or intraperitoneally) into the mammal. It will be understood in the art that the size of the subject and the imaging system used will determine the quantity of imaging moiety needed to produce diagnostic images. In the case of a radioisotope moiety, for a human subject, the quantity of radioactivity injected will normally range from about 5 to 20 millicuries of 99mTc. The labeled antibody or antibody fragment will then preferentially accumulate at the location of cells which contain the specific protein. In vivo tumor imaging is described in S.W. Burchiel et al., "Immunopharmacokinetics of Radiolabeled Antibodies and Their Fragments." (Chapter 13 in Tumor Imaging: The Radiochemical Detection of Cancer, S.W. Burchiel and B. A. Rhodes, eds., Masson Publishing Inc. (1982).)

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Thus, the invention provides a diagnostic method of a disorder, which involves (a) assaying the expression of a polypeptide of the present invention in cells or body fluid of an individual; (b) comparing the level of gene expression with a standard gene expression level, whereby an increase or decrease in the assayed polypeptide gene expression level compared to the standard expression level is indicative of a disorder.

Moreover, polypeptides of the present invention can be used to treat disease. For example, patients can be administered a polypeptide of the present invention in an effort to replace absent or decreased levels of the polypeptide (e.g., insulin), to supplement absent or decreased levels of a different polypeptide (e.g., hemoglobin S for hemoglobin B), to inhibit the activity of a polypeptide (e.g., an oncogene), to activate the activity of a polypeptide (e.g., by binding to a receptor), to reduce the activity of a membrane bound receptor by competing with it for free ligand (e.g., soluble TNF receptors used in reducing inflammation), or to bring about a desired response (e.g., blood vessel growth).

Similarly, antibodies directed to a polypeptide of the present invention can also be used to treat disease. For example, administration of an antibody directed to a polypeptide of the present invention can bind and reduce overproduction of the polypeptide. Similarly, administration of an antibody can activate the polypeptide, such as by binding to a polypeptide bound to a membrane (receptor).

At the very least, the polypeptides of the present invention can be used as molecular weight markers on SDS-PAGE gels or on molecular size gel filtration columns using methods well known to those of skill in the art. Polypeptides can also be used to raise antibodies, which in turn are used to measure protein expression from a recombinant cell, as a way of assessing transformation of the host cell. Moreover, the polypeptides of the present invention can be used to test the following biological activities.

Biological Activities

The polynucleotides and polypeptides of the present invention can be used in assays to test for one or more biological activities. If these polynucleotides and polypeptides do exhibit activity in a particular assay, it is likely that these molecules may be involved in the diseases associated with the biological activity. Thus, the polynucleotides and polypeptides could be used to treat the associated disease.

35 Immune Activity

A polypeptide or polynucleotide of the present invention may be useful in treating deficiencies or disorders of the immune system, by activating or inhibiting the

proliferation, differentiation, or mobilization (chemotaxis) of immune cells. Immune cells develop through a process called hematopoiesis, producing myeloid (platelets, red blood cells, neutrophils, and macrophages) and lymphoid (B and T lymphocytes) cells from pluripotent stem cells. The etiology of these immune deficiencies or disorders may be genetic, somatic, such as cancer or some autoimmune disorders, acquired (e.g., by chemotherapy or toxins), or infectious. Moreover, a polynucleotide or polypeptide of the present invention can be used as a marker or detector of a particular immune system disease or disorder.

A polynucleotide or polypeptide of the present invention may be useful in treating or detecting deficiencies or disorders of hematopoietic cells. A polypeptide or polynucleotide of the present invention could be used to increase differentiation and proliferation of hematopoietic cells, including the pluripotent stem cells, in an effort to treat those disorders associated with a decrease in certain (or many) types hematopoietic cells. Examples of immunologic deficiency syndromes include, but are not limited to: blood protein disorders (e.g. agammaglobulinemia, dysgammaglobulinemia), ataxia telangiectasia, common variable immunodeficiency, Digeorge Syndrome, HIV infection, HTLV-BLV infection, leukocyte adhesion deficiency syndrome, lymphopenia, phagocyte bactericidal dysfunction, severe combined immunodeficiency (SCIDs), Wiskott-Aldrich Disorder, anemia, thrombocytopenia, or hemoglobinuria.

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Moreover, a polypeptide or polynucleotide of the present invention could also be used to modulate hemostatic (the stopping of bleeding) or thrombolytic activity (clot formation). For example, by increasing hemostatic or thrombolytic activity, a polynucleotide or polypeptide of the present invention could be used to treat blood coagulation disorders (e.g., afibrinogenemia, factor deficiencies), blood platelet disorders (e.g. thrombocytopenia), or wounds resulting from trauma, surgery, or other causes. Alternatively, a polynucleotide or polypeptide of the present invention that can decrease hemostatic or thrombolytic activity could be used to inhibit or dissolve clotting. These molecules could be important in the treatment of heart attacks (infarction), strokes, or scarring.

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A polynucleotide or polypeptide of the present invention may also be useful in treating or detecting autoimmune disorders. Many autoimmune disorders result from inappropriate recognition of self as foreign material by immune cells. This inappropriate recognition results in an immune response leading to the destruction of the host tissue. Therefore, the administration of a polypeptide or polynucleotide of the present invention that inhibits an immune response, particularly the proliferation, differentiation, or chemotaxis of T-cells, may be an effective therapy in preventing autoimmune disorders.

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Examples of autoimmune disorders that can be treated or detected by the present invention include, but are not limited to: Addison's Disease, hemolytic anemia, antiphospholipid syndrome, rheumatoid arthritis, dermatitis, allergic encephalomyelitis, glomerulonephritis, Goodpasture's Syndrome, Graves' Disease, Multiple Sclerosis, Myasthenia Gravis, Neuritis, Ophthalmia, Bullous Pemphigoid, Pemphigus, Polyendocrinopathies, Purpura, Reiter's Disease, Stiff-Man Syndrome, Autoimmune Thyroiditis, Systemic Lupus Erythematosus, Autoimmune Pulmonary Inflammation, Guillain-Barre Syndrome, insulin dependent diabetes mellitis, and autoimmune inflammatory eye disease.

Similarly, allergic reactions and conditions, such as asthma (particularly allergic asthma) or other respiratory problems, may also be treated by a polypeptide or polynucleotide of the present invention. Moreover, these molecules can be used to treat anaphylaxis, hypersensitivity to an antigenic molecule, or blood group incompatibility.

A polynucleotide or polypeptide of the present invention may also be used to treat and/or prevent organ rejection or graft-versus-host disease (GVHD). Organ rejection occurs by host immune cell destruction of the transplanted tissue through an immune response. Similarly, an immune response is also involved in GVHD, but, in this case, the foreign transplanted immune cells destroy the host tissues. The administration of a polypeptide or polynucleotide of the present invention that inhibits an immune response, particularly the proliferation, differentiation, or chemotaxis of T-cells, may be an effective therapy in preventing organ rejection or GVHD.

Similarly, a polypeptide or polynucleotide of the present invention may also be used to modulate inflammation. For example, the polypeptide or polynucleotide may inhibit the proliferation and differentiation of cells involved in an inflammatory response. These molecules can be used to treat inflammatory conditions, both chronic and acute conditions, including inflammation associated with infection (e.g., septic shock, sepsis, or systemic inflammatory response syndrome (SIRS)), ischemia-reperfusion injury, endotoxin lethality, arthritis, complement-mediated hyperacute rejection, nephritis, cytokine or chemokine induced lung injury, inflammatory bowel disease, Crohn's disease, or resulting from over production of cytokines (e.g., TNF or IL-1.)

Hyperproliferative Disorders

A polypeptide or polynucleotide can be used to treat or detect hyperproliferative disorders, including neoplasms. A polypeptide or polynucleotide of the present invention may inhibit the proliferation of the disorder through direct or indirect

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interactions. Alternatively, a polypeptide or polynucleotide of the present invention may proliferate other cells which can inhibit the hyperproliferative disorder.

For example, by increasing an immune response, particularly increasing antigenic qualities of the hyperproliferative disorder or by proliferating, differentiating, or mobilizing T-cells, hyperproliferative disorders can be treated. This immune response may be increased by either enhancing an existing immune response, or by initiating a new immune response. Alternatively, decreasing an immune response may also be a method of treating hyperproliferative disorders, such as a chemotherapeutic agent.

Examples of hyperproliferative disorders that can be treated or detected by a polynucleotide or polypeptide of the present invention include, but are not limited to neoplasms located in the: abdomen, bone, breast, digestive system, liver, pancreas, peritoneum, endocrine glands (adrenal, parathyroid, pituitary, testicles, ovary, thymus, thyroid), eye, head and neck, nervous (central and peripheral), lymphatic system, pelvic, skin, soft tissue, spleen, thoracic, and urogenital.

Similarly, other hyperproliferative disorders can also be treated or detected by a polynucleotide or polypeptide of the present invention. Examples of such hyperproliferative disorders include, but are not limited to: hypergammaglobulinemia, lymphoproliferative disorders, paraproteinemias, purpura, sarcoidosis, Sezary Syndrome, Waldenstron's Macroglobulinemia, Gaucher's Disease, histiocytosis, and any other hyperproliferative disease, besides neoplasia, located in an organ system listed above.

Infectious Disease

A polypeptide or polynucleotide of the present invention can be used to treat or detect infectious agents. For example, by increasing the immune response, particularly increasing the proliferation and differentiation of B and/or T cells, infectious diseases may be treated. The immune response may be increased by either enhancing an existing immune response, or by initiating a new immune response. Alternatively, the polypeptide or polynucleotide of the present invention may also directly inhibit the infectious agent, without necessarily eliciting an immune response.

Viruses are one example of an infectious agent that can cause disease or symptoms that can be treated or detected by a polynucleotide or polypeptide of the present invention. Examples of viruses, include, but are not limited to the following DNA and RNA viral families: Arbovirus, Adenoviridae, Arenaviridae, Arterivirus, Birnaviridae, Bunyaviridae, Caliciviridae, Circoviridae, Coronaviridae, Flaviviridae, Hepadnaviridae (Hepatitis), Herpesviridae (such as, Cytomegalovirus, Herpes

Simplex, Herpes Zoster), Mononegavirus (e.g., Paramyxoviridae, Morbillivirus, Rhabdoviridae), Orthomyxoviridae (e.g., Influenza), Papovaviridae, Parvoviridae, Picornaviridae, Poxviridae (such as Smallpox or Vaccinia), Reoviridae (e.g., Rotavirus), Retroviridae (HTLV-I, HTLV-II, Lentivirus), and Togaviridae (e.g., Rubivirus). Viruses falling within these families can cause a variety of diseases or symptoms, including, but not limited to: arthritis, bronchiollitis, encephalitis, eye infections (e.g., conjunctivitis, keratitis), chronic fatigue syndrome, hepatitis (A, B, C, E, Chronic Active, Delta), meningitis, opportunistic infections (e.g., AIDS),

pneumonia, Burkitt's Lymphoma, chickenpox, hemorrhagic fever, Measles, Mumps,

Parainfluenza, Rabies, the common cold, Polio, leukemia, Rubella, sexually
transmitted diseases, skin diseases (e.g., Kaposi's, warts), and viremia. A polypeptide
or polynucleotide of the present invention can be used to treat or detect any of these
symptoms or diseases.

Similarly, bacterial or fungal agents that can cause disease or symptoms and that can be treated or detected by a polynucleotide or polypeptide of the present invention 15 include, but not limited to, the following Gram-Negative and Gram-positive bacterial families and fungi: Actinomycetales (e.g., Corynebacterium, Mycobacterium, Norcardia), Aspergillosis, Bacillaceae (e.g., Anthrax, Clostridium), Bacteroidaceae, Blastomycosis, Bordetella, Borrelia, Brucellosis, Candidiasis, Campylobacter, Coccidioidomycosis, Cryptococcosis, Dermatocycoses, Enterobacteriaceae (Klebsiella, 20 Salmonella, Serratia, Yersinia), Erysipelothrix, Helicobacter, Legionellosis, Leptospirosis, Listeria, Mycoplasmatales, Neisseriaceae (e.g., Acinetobacter, Gonorrhea, Menigococcal), Pasteurellacea Infections (e.g., Actinobacillus, Heamophilus, Pasteurella), Pseudomonas, Rickettsiaceae, Chlamydiaceae, Syphilis, and Staphylococcal. These bacterial or fungal families can cause the following diseases 25 or symptoms, including, but not limited to: bacteremia, endocarditis, eye infections (conjunctivitis, tuberculosis, uveitis), gingivitis, opportunistic infections (e.g., AIDS related infections), paronychia, prosthesis-related infections, Reiter's Disease,

respiratory tract infections, such as Whooping Cough or Empyema, sepsis, Lyme

Disease, Cat-Scratch Disease, Dysentery, Paratyphoid Fever, food poisoning,
Typhoid, pneumonia, Gonorrhea, meningitis, Chlamydia, Syphilis, Diphtheria,
Leprosy, Paratuberculosis, Tuberculosis, Lupus, Botulism, gangrene, tetanus,
impetigo, Rheumatic Fever, Scarlet Fever, sexually transmitted diseases, skin diseases
(e.g., cellulitis, dermatocycoses), toxemia, urinary tract infections, wound infections.

A polypeptide or polynucleotide of the present invention can be used to treat or detect any of these symptoms or diseases.

Moreover, parasitic agents causing disease or symptoms that can be treated or detected by a polynucleotide or polypeptide of the present invention include, but not limited to, the following families: Amebiasis, Babesiosis, Coccidiosis, Cryptosporidiosis, Dientamoebiasis, Dourine, Ectoparasitic, Giardiasis, Helminthiasis, Leishmaniasis, Theileriasis, Toxoplasmosis, Trypanosomiasis, and Trichomonas. These parasites can cause a variety of diseases or symptoms, including, but not limited to: Scabies, Trombiculiasis, eye infections, intestinal disease (e.g., dysentery, giardiasis), liver disease, lung disease, opportunistic infections (e.g., AIDS related), Malaria, pregnancy complications, and toxoplasmosis. A polypeptide or polynucleotide of the present invention can be used to treat or detect any of these symptoms or diseases.

Preferably, treatment using a polypeptide or polynucleotide of the present invention could either be by administering an effective amount of a polypeptide to the patient, or by removing cells from the patient, supplying the cells with a polynucleotide of the present invention, and returning the engineered cells to the patient (ex vivo therapy). Moreover, the polypeptide or polynucleotide of the present invention can be used as an antigen in a vaccine to raise an immune response against infectious disease.

Regeneration

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A polynucleotide or polypeptide of the present invention can be used to differentiate, proliferate, and attract cells, leading to the regeneration of tissues. (See, Science 276:59-87 (1997).) The regeneration of tissues could be used to repair, replace, or protect tissue damaged by congenital defects, trauma (wounds, burns, incisions, or ulcers), age, disease (e.g. osteoporosis, osteocarthritis, periodontal disease, liver failure), surgery, including cosmetic plastic surgery, fibrosis, reperfusion injury, or systemic cytokine damage.

Tissues that could be regenerated using the present invention include organs (e.g., pancreas, liver, intestine, kidney, skin, endothelium), muscle (smooth, skeletal or cardiac), vascular (including vascular endothelium), nervous, hematopoietic, and skeletal (bone, cartilage, tendon, and ligament) tissue. Preferably, regeneration occurs without or decreased scarring. Regeneration also may include angiogenesis.

Moreover, a polynucleotide or polypeptide of the present invention may increase regeneration of tissues difficult to heal. For example, increased tendon/ligament regeneration would quicken recovery time after damage. A polynucleotide or polypeptide of the present invention could also be used prophylactically in an effort to avoid damage. Specific diseases that could be treated include of tendinitis, carpal tunnel syndrome, and other tendon or ligament defects. A further example of tissue

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regeneration of non-healing wounds includes pressure ulcers, ulcers associated with vascular insufficiency, surgical, and traumatic wounds.

Similarly, nerve and brain tissue could also be regenerated by using a polynucleotide or polypeptide of the present invention to proliferate and differentiate nerve cells. Diseases that could be treated using this method include central and peripheral nervous system diseases, neuropathies, or mechanical and traumatic disorders (e.g., spinal cord disorders, head trauma, cerebrovascular disease, and stoke). Specifically, diseases associated with peripheral nerve injuries, peripheral neuropathy (e.g., resulting from chemotherapy or other medical therapies), localized neuropathies, and central nervous system diseases (e.g., Alzheimer's disease, Parkinson's disease, Huntington's disease, amyotrophic lateral sclerosis, and Shy-Drager syndrome), could all be treated using the polynucleotide or polypeptide of the present invention.

15 Chemotaxis

A polynucleotide or polypeptide of the present invention may have chemotaxis activity. A chemotaxic molecule attracts or mobilizes cells (e.g., monocytes, fibroblasts, neutrophils, T-cells, mast cells, eosinophils, epithelial and/or endothelial cells) to a particular site in the body, such as inflammation, infection, or site of hyperproliferation. The mobilized cells can then fight off and/or heal the particular trauma or abnormality.

A polynucleotide or polypeptide of the present invention may increase chemotaxic activity of particular cells. These chemotactic molecules can then be used to treat inflammation, infection, hyperproliferative disorders, or any immune system disorder by increasing the number of cells targeted to a particular location in the body. For example, chemotaxic molecules can be used to treat wounds and other trauma to tissues by attracting immune cells to the injured location. Chemotactic molecules of the present invention can also attract fibroblasts, which can be used to treat wounds.

It is also contemplated that a polynucleotide or polypeptide of the present invention may inhibit chemotactic activity. These molecules could also be used to treat disorders. Thus, a polynucleotide or polypeptide of the present invention could be used as an inhibitor of chemotaxis.

Binding Activity

A polypeptide of the present invention may be used to screen for molecules that bind to the polypeptide or for molecules to which the polypeptide binds. The binding of the polypeptide and the molecule may activate (agonist), increase, inhibit

(antagonist), or decrease activity of the polypeptide or the molecule bound. Examples of such molecules include antibodies, oligonucleotides, proteins (e.g., receptors), or small molecules.

Preferably, the molecule is closely related to the natural ligand of the polypeptide, e.g., a fragment of the ligand, or a natural substrate, a ligand, a structural or functional mimetic. (See, Coligan et al., Current Protocols in Immunology 1(2):Chapter 5 (1991).) Similarly, the molecule can be closely related to the natural receptor to which the polypeptide binds, or at least, a fragment of the receptor capable of being bound by the polypeptide (e.g., active site). In either case, the molecule can be rationally designed using known techniques.

Preferably, the screening for these molecules involves producing appropriate cells which express the polypeptide, either as a secreted protein or on the cell membrane. Preferred cells include cells from mammals, yeast, Drosophila, or *E. coli*. Cells expressing the polypeptide (or cell membrane containing the expressed polypeptide) are then preferably contacted with a test compound potentially containing the molecule to observe binding, stimulation, or inhibition of activity of either the polypeptide or the molecule.

The assay may simply test binding of a candidate compound to the polypeptide, wherein binding is detected by a label, or in an assay involving competition with a labeled competitor. Further, the assay may test whether the candidate compound results in a signal generated by binding to the polypeptide.

Alternatively, the assay can be carried out using cell-free preparations, polypeptide/molecule affixed to a solid support, chemical libraries, or natural product mixtures. The assay may also simply comprise the steps of mixing a candidate compound with a solution containing a polypeptide, measuring polypeptide/molecule activity or binding, and comparing the polypeptide/molecule activity or binding to a standard.

Preferably, an ELISA assay can measure polypeptide level or activity in a sample (e.g., biological sample) using a monoclonal or polyclonal antibody. The antibody can measure polypeptide level or activity by either binding, directly or indirectly, to the polypeptide or by competing with the polypeptide for a substrate.

All of these above assays can be used as diagnostic or prognostic markers. The molecules discovered using these assays can be used to treat disease or to bring about a particular result in a patient (e.g., blood vessel growth) by activating or inhibiting the polypeptide/molecule. Moreover, the assays can discover agents which may inhibit or enhance the production of the polypeptide from suitably manipulated cells or tissues.

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Therefore, the invention includes a method of identifying compounds which bind to a polypeptide of the invention comprising the steps of: (a) incubating a candidate binding compound with a polypeptide of the invention; and (b) determining if binding has occurred. Moreover, the invention includes a method of identifying agonists/antagonists comprising the steps of: (a) incubating a candidate compound with a polypeptide of the invention, (b) assaying a biological activity, and (b) determining if a biological activity of the polypeptide has been altered.

Other Activities

A polypeptide or polynucleotide of the present invention may also increase or decrease the differentiation or proliferation of embryonic stem cells, besides, as discussed above, hematopoietic lineage.

A polypeptide or polynucleotide of the present invention may also be used to modulate mammalian characteristics, such as body height, weight, hair color, eye color, skin, percentage of adipose tissue, pigmentation, size, and shape (e.g., cosmetic surgery). Similarly, a polypeptide or polynucleotide of the present invention may be used to modulate mammalian metabolism affecting catabolism, anabolism, processing, utilization, and storage of energy.

A polypeptide or polynucleotide of the present invention may be used to change a mammal's mental state or physical state by influencing biorhythms, caricadic rhythms, depression (including depressive disorders), tendency for violence, tolerance for pain, reproductive capabilities (preferably by Activin or Inhibin-like activity), hormonal or endocrine levels, appetite, libido, memory, stress, or other cognitive qualities.

A polypeptide or polynucleotide of the present invention may also be used as a food additive or preservative, such as to increase or decrease storage capabilities, fat content, lipid, protein, carbohydrate, vitamins, minerals, cofactors or other nutritional components.

30 Other Preferred Embodiments

Other preferred embodiments of the claimed invention include an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to a sequence of at least about 50 contiguous nucleotides in the nucleotide sequence of SEQ ID NO:X wherein X is any integer as defined in Table 1.

Also preferred is a nucleic acid molecule wherein said sequence of contiguous nucleotides is included in the nucleotide sequence of SEQ ID NO:X in the range of

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positions beginning with the nucleotide at about the position of the 5' Nucleotide of the Clone Sequence and ending with the nucleotide at about the position of the 3' Nucleotide of the Clone Sequence as defined for SEQ ID NO:X in Table 1.

Also preferred is a nucleic acid molecule wherein said sequence of contiguous nucleotides is included in the nucleotide sequence of SEQ ID NO:X in the range of positions beginning with the nucleotide at about the position of the 5' Nucleotide of the Start Codon and ending with the nucleotide at about the position of the 3' Nucleotide of the Clone Sequence as defined for SEQ ID NO:X in Table 1.

Similarly preferred is a nucleic acid molecule wherein said sequence of contiguous nucleotides is included in the nucleotide sequence of SEQ ID NO:X in the range of positions beginning with the nucleotide at about the position of the 5' Nucleotide of the First Amino Acid of the Signal Peptide and ending with the nucleotide at about the position of the 3' Nucleotide of the Clone Sequence as defined for SEQ ID NO:X in Table 1.

Also preferred is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to a sequence of at least about 150 contiguous nucleotides in the nucleotide sequence of SEQ ID NO:X.

Further preferred is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to a sequence of at least about 500 contiguous nucleotides in the nucleotide sequence of SEQ ID NO:X.

A further preferred embodiment is a nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to the nucleotide sequence of SEQ ID NO:X beginning with the nucleotide at about the position of the 5' Nucleotide of the First Amino Acid of the Signal Peptide and ending with the nucleotide at about the position of the 3' Nucleotide of the Clone Sequence as defined for SEQ ID NO:X in Table 1.

A further preferred embodiment is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to the complete nucleotide sequence of SEQ ID NO:X.

Also preferred is an isolated nucleic acid molecule which hybridizes under stringent hybridization conditions to a nucleic acid molecule, wherein said nucleic acid molecule which hybridizes does not hybridize under stringent hybridization conditions to a nucleic acid molecule having a nucleotide sequence consisting of only A residues or of only T residues.

Also preferred is a composition of matter comprising a DNA molecule which comprises a human cDNA clone identified by a cDNA Clone Identifier in Table 1, which DNA molecule is contained in the material deposited with the American Type

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Culture Collection and given the ATCC Deposit Number shown in Table 1 for said cDNA Clone Identifier.

Also preferred is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to a sequence of at least 50 contiguous nucleotides in the nucleotide sequence of a human cDNA clone identified by a cDNA Clone Identifier in Table 1, which DNA molecule is contained in the deposit given the ATCC Deposit Number shown in Table 1.

Also preferred is an isolated nucleic acid molecule, wherein said sequence of at least 50 contiguous nucleotides is included in the nucleotide sequence of the complete open reading frame sequence encoded by said human cDNA clone.

Also preferred is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to sequence of at least 150 contiguous nucleotides in the nucleotide sequence encoded by said human cDNA clone.

A further preferred embodiment is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to sequence of at least 500 contiguous nucleotides in the nucleotide sequence encoded by said human cDNA clone.

A further preferred embodiment is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to the complete nucleotide sequence encoded by said human cDNA clone.

A further preferred embodiment is a method for detecting in a biological sample a nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to a sequence of at least 50 contiguous nucleotides in a sequence selected from the group consisting of: a nucleotide sequence of SEQ ID NO:X wherein X is any integer as defined in Table 1; and a nucleotide sequence encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1; which method comprises a step of comparing a nucleotide sequence of at least one nucleic acid molecule in said sample with a sequence selected from said group and determining whether the sequence of said nucleic acid molecule in said sample is at least 95% identical to said selected sequence.

Also preferred is the above method wherein said step of comparing sequences comprises determining the extent of nucleic acid hybridization between nucleic acid molecules in said sample and a nucleic acid molecule comprising said sequence selected from said group. Similarly, also preferred is the above method wherein said step of comparing sequences is performed by comparing the nucleotide sequence determined from a nucleic acid molecule in said sample with said sequence selected from said group. The nucleic acid molecules can comprise DNA molecules or RNA molecules.

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A further preferred embodiment is a method for identifying the species, tissue or cell type of a biological sample which method comprises a step of detecting nucleic acid molecules in said sample, if any, comprising a nucleotide sequence that is at least 95% identical to a sequence of at least 50 contiguous nucleotides in a sequence selected from the group consisting of: a nucleotide sequence of SEQ ID NO:X wherein X is any integer as defined in Table 1; and a nucleotide sequence encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

The method for identifying the species, tissue or cell type of a biological sample can comprise a step of detecting nucleic acid molecules comprising a nucleotide sequence in a panel of at least two nucleotide sequences, wherein at least one sequence in said panel is at least 95% identical to a sequence of at least 50 contiguous nucleotides in a sequence selected from said group.

Also preferred is a method for diagnosing in a subject a pathological condition associated with abnormal structure or expression of a gene encoding a secreted protein identified in Table 1, which method comprises a step of detecting in a biological sample obtained from said subject nucleic acid molecules, if any, comprising a nucleotide sequence that is at least 95% identical to a sequence of at least 50 contiguous nucleotides in a sequence selected from the group consisting of: a nucleotide sequence of SEQ ID NO:X wherein X is any integer as defined in Table 1; and a nucleotide sequence encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

The method for diagnosing a pathological condition can comprise a step of detecting nucleic acid molecules comprising a nucleotide sequence in a panel of at least two nucleotide sequences, wherein at least one sequence in said panel is at least 95% identical to a sequence of at least 50 contiguous nucleotides in a sequence selected from said group.

Also preferred is a composition of matter comprising isolated nucleic acid molecules wherein the nucleotide sequences of said nucleic acid molecules comprise a panel of at least two nucleotide sequences, wherein at least one sequence in said panel is at least 95% identical to a sequence of at least 50 contiguous nucleotides in a sequence selected from the group consisting of: a nucleotide sequence of SEQ ID NO:X wherein X is any integer as defined in Table 1; and a nucleotide sequence encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1. The nucleic acid molecules can comprise DNA molecules or RNA molecules.

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Also preferred is an isolated polypeptide comprising an amino acid sequence at least 90% identical to a sequence of at least about 10 contiguous amino acids in the amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1.

Also preferred is a polypeptide, wherein said sequence of contiguous amino acids is included in the amino acid sequence of SEQ ID NO:Y in the range of positions beginning with the residue at about the position of the First Amino Acid of the Secreted Portion and ending with the residue at about the Last Amino Acid of the Open Reading Frame as set forth for SEQ ID NO:Y in Table 1.

Also preferred is an isolated polypeptide comprising an amino acid sequence at least 95% identical to a sequence of at least about 30 contiguous amino acids in the amino acid sequence of SEQ ID NO:Y.

Further preferred is an isolated polypeptide comprising an amino acid sequence at least 95% identical to a sequence of at least about 100 contiguous amino acids in the amino acid sequence of SEQ ID NO:Y.

Further preferred is an isolated polypeptide comprising an amino acid sequence at least 95% identical to the complete amino acid sequence of SEQ ID NO:Y.

Further preferred is an isolated polypeptide comprising an amino acid sequence at least 90% identical to a sequence of at least about 10 contiguous amino acids in the complete amino acid sequence of a secreted protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Also preferred is a polypeptide wherein said sequence of contiguous amino acids is included in the amino acid sequence of a secreted portion of the secreted protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Also preferred is an isolated polypeptide comprising an amino acid sequence at least 95% identical to a sequence of at least about 30 contiguous amino acids in the amino acid sequence of the secreted portion of the protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Also preferred is an isolated polypeptide comprising an amino acid sequence at least 95% identical to a sequence of at least about 100 contiguous amino acids in the amino acid sequence of the secreted portion of the protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

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Also preferred is an isolated polypeptide comprising an amino acid sequence at least 95% identical to the amino acid sequence of the secreted portion of the protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Further preferred is an isolated antibody which binds specifically to a polypeptide comprising an amino acid sequence that is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Further preferred is a method for detecting in a biological sample a polypeptide comprising an amino acid sequence which is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1; which method comprises a step of comparing an amino acid sequence of at least one polypeptide molecule in said sample with a sequence selected from said group and determining whether the sequence of said polypeptide molecule in said sample is at least 90% identical to said sequence of at least 10 contiguous amino acids.

Also preferred is the above method wherein said step of comparing an amino acid sequence of at least one polypeptide molecule in said sample with a sequence selected from said group comprises determining the extent of specific binding of polypeptides in said sample to an antibody which binds specifically to a polypeptide comprising an amino acid sequence that is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Also preferred is the above method wherein said step of comparing sequences is performed by comparing the amino acid sequence determined from a polypeptide molecule in said sample with said sequence selected from said group.

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Also preferred is a method for identifying the species, tissue or cell type of a biological sample which method comprises a step of detecting polypeptide molecules in said sample, if any, comprising an amino acid sequence that is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a secreted protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Also preferred is the above method for identifying the species, tissue or cell type of a biological sample, which method comprises a step of detecting polypeptide molecules comprising an amino acid sequence in a panel of at least two amino acid sequences, wherein at least one sequence in said panel is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the above group.

Also preferred is a method for diagnosing in a subject a pathological condition associated with abnormal structure or expression of a gene encoding a secreted protein identified in Table 1, which method comprises a step of detecting in a biological sample obtained from said subject polypeptide molecules comprising an amino acid sequence in a panel of at least two amino acid sequences, wherein at least one sequence in said panel is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a secreted protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

In any of these methods, the step of detecting said polypeptide molecules includes using an antibody.

Also preferred is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to a nucleotide sequence encoding a polypeptide wherein said polypeptide comprises an amino acid sequence that is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a secreted protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

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Also preferred is an isolated nucleic acid molecule, wherein said nucleotide sequence encoding a polypeptide has been optimized for expression of said polypeptide in a prokaryotic host.

Also preferred is an isolated nucleic acid molecule, wherein said polypeptide comprises an amino acid sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a secreted protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Further preferred is a method of making a recombinant vector comprising inserting any of the above isolated nucleic acid molecule into a vector. Also preferred is the recombinant vector produced by this method. Also preferred is a method of making a recombinant host cell comprising introducing the vector into a host cell, as well as the recombinant host cell produced by this method.

Also preferred is a method of making an isolated polypeptide comprising culturing this recombinant host cell under conditions such that said polypeptide is expressed and recovering said polypeptide. Also preferred is this method of making an isolated polypeptide, wherein said recombinant host cell is a eukaryotic cell and said polypeptide is a secreted portion of a human secreted protein comprising an amino acid sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y beginning with the residue at the position of the First Amino Acid of the Secreted Portion of SEQ ID NO:Y wherein Y is an integer set forth in Table 1 and said position of the First Amino Acid of the Secreted Portion of SEQ ID NO:Y is defined in Table 1; and an amino acid sequence of a secreted portion of a protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1. The isolated polypeptide produced by this method is also preferred.

Also preferred is a method of treatment of an individual in need of an increased level of a secreted protein activity, which method comprises administering to such an individual a pharmaceutical composition comprising an amount of an isolated polypeptide, polynucleotide, or antibody of the claimed invention effective to increase the level of said protein activity in said individual.

Having generally described the invention, the same will be more readily understood by reference to the following examples, which are provided by way of illustration and are not intended as limiting.

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Examples

Example 1: Isolation of a Selected cDNA Clone From the Deposited Sample

Each cDNA clone in a cited ATCC deposit is contained in a plasmid vector. Table 1 identifies the vectors used to construct the cDNA library from which each clone was isolated. In many cases, the vector used to construct the library is a phage vector from which a plasmid has been excised. The table immediately below correlates the related plasmid for each phage vector used in constructing the cDNA library. For example, where a particular clone is identified in Table 1 as being isolated in the vector "Lambda Zap," the corresponding deposited clone is in "pBluescript."

	Vector Used to Construct Library	Corresponding Deposited Plasmid		
	Lambda Zap	pBluescript (pBS)		
	Uni-Zap XR	pBluescript (pBS)		
15	Zap Express	pBK		
	lafmid BA	plafmid BA		
	pSport1	pSport1		
	pCMVSport 2.0	pCMVSport 2.0		
	pCMVSport 3.0	pCMVSport 3.0		
20	pCR®2.1	pCR®2.1		

Vectors Lambda Zap (U.S. Patent Nos. 5,128,256 and 5,286,636), Uni-Zap XR (U.S. Patent Nos. 5,128, 256 and 5,286,636), Zap Express (U.S. Patent Nos. 5,128,256 and 5,286,636), pBluescript (pBS) (Short, J. M. et al., Nucleic Acids Res. 16:7583-7600 (1988); Alting-Mees, M. A. and Short, J. M., Nucleic Acids Res.

- 17:9494 (1989)) and pBK (Alting-Mees, M. A. et al., Strategies 5:58-61 (1992)) are commercially available from Stratagene Cloning Systems, Inc., 11011 N. Torrey Pines Road, La Jolla, CA, 92037. pBS contains an ampicillin resistance gene and pBK contains a neomycin resistance gene. Both can be transformed into E. coli strain XL-1 Blue, also available from Stratagene. pBS comes in 4 forms SK+, SK-, KS+ and KS.
- The S and K refers to the orientation of the polylinker to the T7 and T3 primer sequences which flank the polylinker region ("S" is for SacI and "K" is for KpnI which are the first sites on each respective end of the linker). "+" or "-" refer to the orientation of the f1 origin of replication ("ori"), such that in one orientation, single stranded rescue initiated from the f1 ori generates sense strand DNA and in the other, antisense.

Vectors pSport1, pCMVSport 2.0 and pCMVSport 3.0, were obtained from Life Technologies, Inc., P. O. Box 6009, Gaithersburg, MD 20897. All Sport vectors contain an ampicillin resistance gene and may be transformed into E. coli strain

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DH10B, also available from Life Technologies. (See, for instance, Gruber, C. E., et al., Focus 15:59 (1993).) Vector lafmid BA (Bento Soares, Columbia University, NY) contains an ampicillin resistance gene and can be transformed into E. coli strain XL-1 Blue. Vector pCR®2.1, which is available from Invitrogen, 1600 Faraday Avenue, Carlsbad, CA 92008, contains an ampicillin resistance gene and may be transformed into E. coli strain DH10B, available from Life Technologies. (See, for instance, Clark, J. M., Nuc. Acids Res. 16:9677-9686 (1988) and Mead, D. et al., Bio/Technology 9: (1991).) Preferably, a polynucleotide of the present invention does not comprise the phage vector sequences identified for the particular clone in Table 1, as well as the corresponding plasmid vector sequences designated above.

The deposited material in the sample assigned the ATCC Deposit Number cited in Table 1 for any given cDNA clone also may contain one or more additional plasmids, each comprising a cDNA clone different from that given clone. Thus, deposits sharing the same ATCC Deposit Number contain at least a plasmid for each cDNA clone identified in Table 1. Typically, each ATCC deposit sample cited in Table 1 comprises a mixture of approximately equal amounts (by weight) of about 50 plasmid DNAs, each containing a different cDNA clone; but such a deposit sample may include plasmids for more or less than 50 cDNA clones, up to about 500 cDNA clones.

Two approaches can be used to isolate a particular clone from the deposited sample of plasmid DNAs cited for that clone in Table 1. First, a plasmid is directly isolated by screening the clones using a polynucleotide probe corresponding to SEQ ID NO:X.

Particularly, a specific polynucleotide with 30-40 nucleotides is synthesized

using an Applied Biosystems DNA synthesizer according to the sequence reported.

The oligonucleotide is labeled, for instance, with ³²P-γ-ATP using T4 polynucleotide kinase and purified according to routine methods. (E.g., Maniatis et al., Molecular Cloning: A Laboratory Manual, Cold Spring Harbor Press, Cold Spring, NY (1982).)

The plasmid mixture is transformed into a suitable host, as indicated above (such as XL-1 Blue (Stratagene)) using techniques known to those of skill in the art, such as those provided by the vector supplier or in related publications or patents cited above. The transformants are plated on 1.5% agar plates (containing the appropriate selection agent, e.g., ampicillin) to a density of about 150 transformants (colonies) per plate. These plates are screened using Nylon membranes according to routine methods for bacterial colony screening (e.g., Sambrook et al., Molecular Cloning: A Laboratory Manual, 2nd Edit., (1989), Cold Spring Harbor Laboratory Press, pages 1.93 to

1.104), or other techniques known to those of skill in the art.

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Alternatively, two primers of 17-20 nucleotides derived from both ends of the SEQ ID NO:X (i.e., within the region of SEQ ID NO:X bounded by the 5' NT and the 3' NT of the clone defined in Table 1) are synthesized and used to amplify the desired cDNA using the deposited cDNA plasmid as a template. The polymerase chain reaction is carried out under routine conditions, for instance, in 25 µl of reaction mixture with 0.5 ug of the above cDNA template. A convenient reaction mixture is 1.5-5 mM MgCl₂, 0.01% (w/v) gelatin, 20 µM each of dATP, dCTP, dGTP, dTTP, 25 pmol of each primer and 0.25 Unit of Taq polymerase. Thirty five cycles of PCR (denaturation at 94°C for 1 min; annealing at 55°C for 1 min; elongation at 72°C for 1 min) are performed with a Perkin-Elmer Cetus automated thermal cycler. The amplified product is analyzed by agarose gel electrophoresis and the DNA band with expected molecular weight is excised and purified. The PCR product is verified to be the selected sequence by subcloning and sequencing the DNA product.

Several methods are available for the identification of the 5' or 3' non-coding portions of a gene which may not be present in the deposited clone. These methods include but are not limited to, filter probing, clone enrichment using specific probes, and protocols similar or identical to 5' and 3' "RACE" protocols which are well known in the art. For instance, a method similar to 5' RACE is available for generating the missing 5' end of a desired full-length transcript. (Fromont-Racine et al., Nucleic Acids Res. 21(7):1683-1684 (1993).)

Briefly, a specific RNA oligonucleotide is ligated to the 5' ends of a population of RNA presumably containing full-length gene RNA transcripts. A primer set containing a primer specific to the ligated RNA oligonucleotide and a primer specific to a known sequence of the gene of interest is used to PCR amplify the 5' portion of the desired full-length gene. This amplified product may then be sequenced and used to generate the full length gene.

This above method starts with total RNA isolated from the desired source, although poly-A+ RNA can be used. The RNA preparation can then be treated with phosphatase if necessary to eliminate 5' phosphate groups on degraded or damaged RNA which may interfere with the later RNA ligase step. The phosphatase should then be inactivated and the RNA treated with tobacco acid pyrophosphatase in order to remove the cap structure present at the 5' ends of messenger RNAs. This reaction leaves a 5' phosphate group at the 5' end of the cap cleaved RNA which can then be ligated to an RNA oligonucleotide using T4 RNA ligase.

This modified RNA preparation is used as a template for first strand cDNA synthesis using a gene specific oligonucleotide. The first strand synthesis reaction is

used as a template for PCR amplification of the desired 5' end using a primer specific to the ligated RNA oligonucleotide and a primer specific to the known sequence of the gene of interest. The resultant product is then sequenced and analyzed to confirm that the 5' end sequence belongs to the desired gene.

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Example 2: Isolation of Genomic Clones Corresponding to a **Polynucleotide**

A human genomic P1 library (Genomic Systems, Inc.) is screened by PCR using primers selected for the cDNA sequence corresponding to SEQ ID NO:X., according to the method described in Example 1. (See also, Sambrook.)

Example 3: Tissue Distribution of Polypeptide

Tissue distribution of mRNA expression of polynucleotides of the present invention is determined using protocols for Northern blot analysis, described by, among others, Sambrook et al. For example, a cDNA probe produced by the method described in Example 1 is labeled with P32 using the rediprimeTM DNA labeling system (Amersham Life Science), according to manufacturer's instructions. After labeling, the probe is purified using CHROMA SPIN-100™ column (Clontech Laboratories, Inc.), according to manufacturer's protocol number PT1200-1. The purified labeled probe is then used to examine various human tissues for mRNA expression.

Multiple Tissue Northern (MTN) blots containing various human tissues (H) or human immune system tissues (IM) (Clontech) are examined with the labeled probe using ExpressHybTM hybridization solution (Clontech) according to manufacturer's protocol number PT1190-1. Following hybridization and washing, the blots are mounted and exposed to film at -70°C overnight, and the films developed according to standard procedures.

Example 4: Chromosomal Mapping of the Polynucleotides

An oligonucleotide primer set is designed according to the sequence at the 5' end of SEQ ID NO:X. This primer preferably spans about 100 nucleotides. This primer set is then used in a polymerase chain reaction under the following set of conditions: 30 seconds, 95°C; 1 minute, 56°C; 1 minute, 70°C. This cycle is repeated 32 times followed by one 5 minute cycle at 70°C. Human, mouse, and hamster DNA is used as template in addition to a somatic cell hybrid panel containing individual chromosomes or chromosome fragments (Bios, Inc). The reactions is analyzed on

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either 8% polyacrylamide gels or 3.5 % agarose gels. Chromosome mapping is determined by the presence of an approximately 100 bp PCR fragment in the particular somatic cell hybrid.

5 Example 5: Bacterial Expression of a Polypeptide

A polynucleotide encoding a polypeptide of the present invention is amplified using PCR oligonucleotide primers corresponding to the 5' and 3' ends of the DNA sequence, as outlined in Example 1, to synthesize insertion fragments. The primers used to amplify the cDNA insert should preferably contain restriction sites, such as BamHI and XbaI, at the 5' end of the primers in order to clone the amplified product into the expression vector. For example, BamHI and XbaI correspond to the restriction enzyme sites on the bacterial expression vector pQE-9. (Qiagen, Inc., Chatsworth, CA). This plasmid vector encodes antibiotic resistance (Amp^r), a bacterial origin of replication (ori), an IPTG-regulatable promoter/operator (P/O), a ribosome binding site (RBS), a 6-histidine tag (6-His), and restriction enzyme cloning sites.

The pQE-9 vector is digested with BamHI and XbaI and the amplified fragment is ligated into the pQE-9 vector maintaining the reading frame initiated at the bacterial RBS. The ligation mixture is then used to transform the E. coli strain M15/rep4 (Qiagen, Inc.) which contains multiple copies of the plasmid pREP4, which expresses the lacI repressor and also confers kanamycin resistance (Kan^r). Transformants are identified by their ability to grow on LB plates and ampicillin/kanamycin resistant colonies are selected. Plasmid DNA is isolated and confirmed by restriction analysis.

Clones containing the desired constructs are grown overnight (O/N) in liquid culture in LB media supplemented with both Amp (100 ug/ml) and Kan (25 ug/ml). The O/N culture is used to inoculate a large culture at a ratio of 1:100 to 1:250. The cells are grown to an optical density 600 (O.D.⁶⁰⁰) of between 0.4 and 0.6. IPTG (Isopropyl-B-D-thiogalacto pyranoside) is then added to a final concentration of 1 mM. IPTG induces by inactivating the lacI repressor, clearing the P/O leading to increased gene expression.

Cells are grown for an extra 3 to 4 hours. Cells are then harvested by centrifugation (20 mins at 6000Xg). The cell pellet is solubilized in the chaotropic agent 6 Molar Guanidine HCl by stirring for 3-4 hours at 4°C. The cell debris is removed by centrifugation, and the supernatant containing the polypeptide is loaded onto a nickel-nitrilo-tri-acetic acid ("Ni-NTA") affinity resin column (available from QIAGEN, Inc., *supra*). Proteins with a 6 x His tag bind to the Ni-NTA resin with high

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affinity and can be purified in a simple one-step procedure (for details see: The QIAexpressionist (1995) QIAGEN, Inc., *supra*).

Briefly, the supernatant is loaded onto the column in 6 M guanidine-HCl, pH 8, the column is first washed with 10 volumes of 6 M guanidine-HCl, pH 8, then washed with 10 volumes of 6 M guanidine-HCl pH 6, and finally the polypeptide is eluted with 6 M guanidine-HCl, pH 5.

The purified protein is then renatured by dialyzing it against phosphate-buffered saline (PBS) or 50 mM Na-acetate, pH 6 buffer plus 200 mM NaCl. Alternatively, the protein can be successfully refolded while immobilized on the Ni-NTA column. The recommended conditions are as follows: renature using a linear 6M-1M urea gradient in 500 mM NaCl, 20% glycerol, 20 mM Tris/HCl pH 7.4, containing protease inhibitors. The renaturation should be performed over a period of 1.5 hours or more. After renaturation the proteins are eluted by the addition of 250 mM immidazole. Immidazole is removed by a final dialyzing step against PBS or 50 mM sodium acetate pH 6 buffer plus 200 mM NaCl. The purified protein is stored at 4°C or frozen at -80°C.

In addition to the above expression vector, the present invention further includes an expression vector comprising phage operator and promoter elements operatively linked to a polynucleotide of the present invention, called pHE4a. (ATCC Accession Number 209645, deposited on February 25, 1998.) This vector contains: 1) a neomycinphosphotransferase gene as a selection marker, 2) an E. coli origin of replication, 3) a T5 phage promoter sequence, 4) two lac operator sequences, 5) a Shine-Delgarno sequence, and 6) the lactose operon repressor gene (lacIq). The origin of replication (oriC) is derived from pUC19 (LTI, Gaithersburg, MD). The promoter sequence and operator sequences are made synthetically.

DNA can be inserted into the pHEa by restricting the vector with NdeI and XbaI, BamHI, XhoI, or Asp718, running the restricted product on a gel, and isolating the larger fragment (the stuffer fragment should be about 310 base pairs). The DNA insert is generated according to the PCR protocol described in Example 1, using PCR primers having restriction sites for NdeI (5' primer) and XbaI, BamHI, XhoI, or Asp718 (3' primer). The PCR insert is gel purified and restricted with compatible enzymes. The insert and vector are ligated according to standard protocols.

The engineered vector could easily be substituted in the above protocol to express protein in a bacterial system.

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Example 6: Purification of a Polypeptide from an Inclusion Body

The following alternative method can be used to purify a polypeptide expressed in $E \, coli$ when it is present in the form of inclusion bodies. Unless otherwise specified, all of the following steps are conducted at 4-10°C.

Upon completion of the production phase of the *E. coli* fermentation, the cell culture is cooled to 4-10°C and the cells harvested by continuous centrifugation at 15,000 rpm (Heraeus Sepatech). On the basis of the expected yield of protein per unit weight of cell paste and the amount of purified protein required, an appropriate amount of cell paste, by weight, is suspended in a buffer solution containing 100 mM Tris, 50 mM EDTA, pH 7.4. The cells are dispersed to a homogeneous suspension using a high shear mixer.

The cells are then lysed by passing the solution through a microfluidizer (Microfuidics, Corp. or APV Gaulin, Inc.) twice at 4000-6000 psi. The homogenate is then mixed with NaCl solution to a final concentration of 0.5 M NaCl, followed by centrifugation at 7000 xg for 15 min. The resultant pellet is washed again using 0.5M NaCl, 100 mM Tris, 50 mM EDTA, pH 7.4.

The resulting washed inclusion bodies are solubilized with 1.5 M guanidine hydrochloride (GuHCl) for 2-4 hours. After 7000 xg centrifugation for 15 min., the pellet is discarded and the polypeptide containing supernatant is incubated at 4°C overnight to allow further GuHCl extraction.

Following high speed centrifugation (30,000 xg) to remove insoluble particles, the GuHCl solubilized protein is refolded by quickly mixing the GuHCl extract with 20 volumes of buffer containing 50 mM sodium, pH 4.5, 150 mM NaCl, 2 mM EDTA by vigorous stirring. The refolded diluted protein solution is kept at 4°C without mixing for 12 hours prior to further purification steps.

To clarify the refolded polypeptide solution, a previously prepared tangential filtration unit equipped with 0.16 µm membrane filter with appropriate surface area (e.g., Filtron), equilibrated with 40 mM sodium acetate, pH 6.0 is employed. The filtered sample is loaded onto a cation exchange resin (e.g., Poros HS-50, Perseptive Biosystems). The column is washed with 40 mM sodium acetate, pH 6.0 and eluted with 250 mM, 500 mM, 1000 mM, and 1500 mM NaCl in the same buffer, in a stepwise manner. The absorbance at 280 nm of the effluent is continuously monitored. Fractions are collected and further analyzed by SDS-PAGE.

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Fractions containing the polypeptide are then pooled and mixed with 4 volumes of water. The diluted sample is then loaded onto a previously prepared set of tandem columns of strong anion (Poros HQ-50, Perseptive Biosystems) and weak anion (Poros CM-20, Perseptive Biosystems) exchange resins. The columns are equilibrated with 40 mM sodium acetate, pH 6.0. Both columns are washed with 40 mM sodium acetate, pH 6.0, 200 mM NaCl. The CM-20 column is then eluted using a 10 column volume linear gradient ranging from 0.2 M NaCl, 50 mM sodium acetate, pH 6.0 to 1.0 M NaCl, 50 mM sodium acetate, pH 6.5. Fractions are collected under constant A₂₈₀ monitoring of the effluent. Fractions containing the polypeptide (determined, for instance, by 16% SDS-PAGE) are then pooled.

The resultant polypeptide should exhibit greater than 95% purity after the above refolding and purification steps. No major contaminant bands should be observed from Commassie blue stained 16% SDS-PAGE gel when 5 μ g of purified protein is loaded. The purified protein can also be tested for endotoxin/LPS contamination, and typically the LPS content is less than 0.1 ng/ml according to LAL assays.

Example 7: Cloning and Expression of a Polypeptide in a Baculovirus Expression System

In this example, the plasmid shuttle vector pA2 is used to insert a polynucleotide into a baculovirus to express a polypeptide. This expression vector contains the strong polyhedrin promoter of the *Autographa californica* nuclear polyhedrosis virus (AcMNPV) followed by convenient restriction sites such as BamHI, Xba I and Asp718. The polyadenylation site of the simian virus 40 ("SV40") is used for efficient polyadenylation. For easy selection of recombinant virus, the plasmid contains the beta-galactosidase gene from *E. coli* under control of a weak Drosophila promoter in the same orientation, followed by the polyadenylation signal of the polyhedrin gene. The inserted genes are flanked on both sides by viral sequences for cell-mediated homologous recombination with wild-type viral DNA to generate a viable virus that express the cloned polynucleotide.

Many other baculovirus vectors can be used in place of the vector above, such as pAc373, pVL941, and pAcIM1, as one skilled in the art would readily appreciate, as long as the construct provides appropriately located signals for transcription, translation, secretion and the like, including a signal peptide and an in-frame AUG as required. Such vectors are described, for instance, in Luckow et al., Virology 170:31-39 (1989).

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Specifically, the cDNA sequence contained in the deposited clone, including the AUG initiation codon and the naturally associated leader sequence identified in Table 1, is amplified using the PCR protocol described in Example 1. If the naturally occurring signal sequence is used to produce the secreted protein, the pA2 vector does not need a second signal peptide. Alternatively, the vector can be modified (pA2 GP) to include a baculovirus leader sequence, using the standard methods described in Summers et al., "A Manual of Methods for Baculovirus Vectors and Insect Cell Culture Procedures," Texas Agricultural Experimental Station Bulletin No. 1555 (1987).

The amplified fragment is isolated from a 1% agarose gel using a commercially available kit ("Geneclean," BIO 101 Inc., La Jolla, Ca.). The fragment then is digested with appropriate restriction enzymes and again purified on a 1% agarose gel.

The plasmid is digested with the corresponding restriction enzymes and optionally, can be dephosphorylated using calf intestinal phosphatase, using routine procedures known in the art. The DNA is then isolated from a 1% agarose gel using a commercially available kit ("Geneclean" BIO 101 Inc., La Jolla, Ca.).

The fragment and the dephosphorylated plasmid are ligated together with T4 DNA ligase. *E. coli* HB101 or other suitable *E. coli* hosts such as XL-1 Blue (Stratagene Cloning Systems, La Jolla, CA) cells are transformed with the ligation mixture and spread on culture plates. Bacteria containing the plasmid are identified by digesting DNA from individual colonies and analyzing the digestion product by gel electrophoresis. The sequence of the cloned fragment is confirmed by DNA sequencing.

Five μg of a plasmid containing the polynucleotide is co-transfected with 1.0 μg of a commercially available linearized baculovirus DNA ("BaculoGold™ baculovirus DNA", Pharmingen, San Diego, CA), using the lipofection method described by Felgner et al., Proc. Natl. Acad. Sci. USA 84:7413-7417 (1987). One μg of BaculoGold™ virus DNA and 5 μg of the plasmid are mixed in a sterile well of a microtiter plate containing 50 μl of serum-free Grace's medium (Life Technologies Inc., Gaithersburg, MD). Afterwards, 10 μl Lipofectin plus 90 μl Grace's medium are added, mixed and incubated for 15 minutes at room temperature. Then the transfection mixture is added drop-wise to Sf9 insect cells (ATCC CRL 1711) seeded in a 35 mm tissue culture plate with 1 ml Grace's medium without serum. The plate is then incubated for 5 hours at 27° C. The transfection solution is then removed from the plate and 1 ml of Grace's insect medium supplemented with 10% fetal calf serum is added. Cultivation is then continued at 27° C for four days.

After four days the supernatant is collected and a plaque assay is performed, as described by Summers and Smith, *supra*. An agarose gel with "Blue Gal" (Life

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Technologies Inc., Gaithersburg) is used to allow easy identification and isolation of gal-expressing clones, which produce blue-stained plaques. (A detailed description of a "plaque assay" of this type can also be found in the user's guide for insect cell culture and baculovirology distributed by Life Technologies Inc., Gaithersburg, page 9-10.) After appropriate incubation, blue stained plaques are picked with the tip of a micropipettor (e.g., Eppendorf). The agar containing the recombinant viruses is then resuspended in a microcentrifuge tube containing 200 µl of Grace's medium and the suspension containing the recombinant baculovirus is used to infect Sf9 cells seeded in 35 mm dishes. Four days later the supernatants of these culture dishes are harvested and then they are stored at 4° C.

To verify the expression of the polypeptide, Sf9 cells are grown in Grace's medium supplemented with 10% heat-inactivated FBS. The cells are infected with the recombinant baculovirus containing the polynucleotide at a multiplicity of infection ("MOI") of about 2. If radiolabeled proteins are desired, 6 hours later the medium is removed and is replaced with SF900 II medium minus methionine and cysteine (available from Life Technologies Inc., Rockville, MD). After 42 hours, 5 μ Ci of ³⁵S-methionine and 5 μ Ci ³⁵S-cysteine (available from Amersham) are added. The cells are further incubated for 16 hours and then are harvested by centrifugation. The proteins in the supernatant as well as the intracellular proteins are analyzed by SDS-PAGE followed by autoradiography (if radiolabeled).

Microsequencing of the amino acid sequence of the amino terminus of purified protein may be used to determine the amino terminal sequence of the produced protein.

Example 8: Expression of a Polypeptide in Mammalian Cells

The polypeptide of the present invention can be expressed in a mammalian cell.

A typical mammalian expression vector contains a promoter element, which mediates the initiation of transcription of mRNA, a protein coding sequence, and signals required for the termination of transcription and polyadenylation of the transcript. Additional elements include enhancers, Kozak sequences and intervening sequences flanked by donor and acceptor sites for RNA splicing. Highly efficient transcription is achieved with the early and late promoters from SV40, the long terminal repeats (LTRs) from Retroviruses, e.g., RSV, HTLVI, HIVI and the early promoter of the cytomegalovirus (CMV). However, cellular elements can also be used (e.g., the human actin promoter).

Suitable expression vectors for use in practicing the present invention include, for example, vectors such as pSVL and pMSG (Pharmacia, Uppsala, Sweden), pRSVcat (ATCC 37152), pSV2dhfr (ATCC 37146), pBC12MI (ATCC 67109), pCMVSport 2.0, and pCMVSport 3.0. Mammalian host cells that could be used

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include, human Hela, 293, H9 and Jurkat cells, mouse NIH3T3 and C127 cells, Cos 1, Cos 7 and CV1, quail QC1-3 cells, mouse L cells and Chinese hamster ovary (CHO) cells.

Alternatively, the polypeptide can be expressed in stable cell lines containing the polynucleotide integrated into a chromosome. The co-transfection with a selectable marker such as dhfr, gpt, neomycin, hygromycin allows the identification and isolation of the transfected cells.

The transfected gene can also be amplified to express large amounts of the encoded protein. The DHFR (dihydrofolate reductase) marker is useful in developing cell lines that carry several hundred or even several thousand copies of the gene of interest. (See, e.g., Alt, F. W., et al., J. Biol. Chem. 253:1357-1370 (1978); Hamlin, J. L. and Ma, C., Biochem. et Biophys. Acta, 1097:107-143 (1990); Page, M. J. and Sydenham, M. A., Biotechnology 9:64-68 (1991).) Another useful selection marker is the enzyme glutamine synthase (GS) (Murphy et al., Biochem J. 227:277-279 (1991); Bebbington et al., Bio/Technology 10:169-175 (1992). Using these markers, the mammalian cells are grown in selective medium and the cells with the highest resistance are selected. These cell lines contain the amplified gene(s) integrated into a chromosome. Chinese hamster ovary (CHO) and NSO cells are often used for the production of proteins.

Derivatives of the plasmid pSV2-dhfr (ATCC Accession No. 37146), the expression vectors pC4 (ATCC Accession No. 209646) and pC6 (ATCC Accession No.209647) contain the strong promoter (LTR) of the Rous Sarcoma Virus (Cullen et al., Molecular and Cellular Biology, 438-447 (March, 1985)) plus a fragment of the CMV-enhancer (Boshart et al., Cell 41:521-530 (1985).) Multiple cloning sites, e.g., with the restriction enzyme cleavage sites BamHI, XbaI and Asp718, facilitate the cloning of the gene of interest. The vectors also contain the 3' intron, the polyadenylation and termination signal of the rat preproinsulin gene, and the mouse DHFR gene under control of the SV40 early promoter.

Specifically, the plasmid pC6, for example, is digested with appropriate restriction enzymes and then dephosphorylated using calf intestinal phosphates by procedures known in the art. The vector is then isolated from a 1% agarose gel.

A polynucleotide of the present invention is amplified according to the protocol outlined in Example 1. If the naturally occurring signal sequence is used to produce the secreted protein, the vector does not need a second signal peptide. Alternatively, if the naturally occurring signal sequence is not used, the vector can be modified to include a heterologous signal sequence. (See, e.g., WO 96/34891.)

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The amplified fragment is isolated from a 1% agarose gel using a commercially available kit ("Geneclean," BIO 101 Inc., La Jolla, Ca.). The fragment then is digested with appropriate restriction enzymes and again purified on a 1% agarose gel.

The amplified fragment is then digested with the same restriction enzyme and purified on a 1% agarose gel. The isolated fragment and the dephosphorylated vector are then ligated with T4 DNA ligase. *E. coli* HB101 or XL-1 Blue cells are then transformed and bacteria are identified that contain the fragment inserted into plasmid pC6 using, for instance, restriction enzyme analysis.

Chinese hamster ovary cells lacking an active DHFR gene is used for transfection. Five µg of the expression plasmid pC6 is cotransfected with 0.5 µg of the plasmid pSVneo using lipofectin (Felgner et al., supra). The plasmid pSV2-neo contains a dominant selectable marker, the neo gene from Tn5 encoding an enzyme that confers resistance to a group of antibiotics including G418. The cells are seeded in alpha minus MEM supplemented with 1 mg/ml G418. After 2 days, the cells are trypsinized and seeded in hybridoma cloning plates (Greiner, Germany) in alpha minus MEM supplemented with 10, 25, or 50 ng/ml of metothrexate plus 1 mg/ml G418. After about 10-14 days single clones are trypsinized and then seeded in 6-well petri dishes or 10 ml flasks using different concentrations of methotrexate (50 nM, 100 nM, 200 nM, 400 nM, 800 nM). Clones growing at the highest concentrations of methotrexate are then transferred to new 6-well plates containing even higher concentrations of methotrexate (1 μM , 2 μM , 5 μM , 10 mM, 20 mM). The same procedure is repeated until clones are obtained which grow at a concentration of 100 -200 µM. Expression of the desired gene product is analyzed, for instance, by SDS-PAGE and Western blot or by reversed phase HPLC analysis.

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Example 9: Protein Fusions

These fusion proteins can be used for a variety of applications. For example, fusion of the present polypeptides to His-tag, HA-tag, protein A, IgG domains, and maltose binding protein facilitates purification. (See Example 5; see also EP A 394,827; Traunecker, et al., Nature 331:84-86 (1988).) Similarly, fusion to IgG-1, IgG-3, and albumin increases the halflife time in vivo. Nuclear localization signals fused to the polypeptides of the present invention can target the protein to a specific subcellular localization, while covalent heterodimer or homodimers can increase or decrease the activity of a fusion protein. Fusion proteins can also create chimeric molecules having more than one function. Finally, fusion proteins can increase solubility and/or stability of the fused protein compared to the non-fused protein. All of the types of fusion

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proteins described above can be made by modifying the following protocol, which outlines the fusion of a polypeptide to an IgG molecule, or the protocol described in Example 5.

Briefly, the human Fc portion of the IgG molecule can be PCR amplified, using primers that span the 5' and 3' ends of the sequence described below. These primers also should have convenient restriction enzyme sites that will facilitate cloning into an expression vector, preferably a mammalian expression vector.

For example, if pC4 (Accession No. 209646) is used, the human Fc portion can be ligated into the BamHI cloning site. Note that the 3' BamHI site should be destroyed. Next, the vector containing the human Fc portion is re-restricted with BamHI, linearizing the vector, and a polynucleotide of the present invention, isolated by the PCR protocol described in Example 1, is ligated into this BamHI site. Note that the polynucleotide is cloned without a stop codon, otherwise a fusion protein will not be produced.

If the naturally occurring signal sequence is used to produce the secreted protein, pC4 does not need a second signal peptide. Alternatively, if the naturally occurring signal sequence is not used, the vector can be modified to include a heterologous signal sequence. (See, e.g., WO 96/34891.)

20 Human IgG Fc region:

GGGATCCGGAGCCCAAATCTTCTGACAAAACTCACACATGCCCACCGTGCC CAGCACCTGAATTCGAGGGTGCACCGTCAGTCTTCCTCTTCCCCCCAAAACC CAAGGACACCCTCATGATCTCCCGGACTCCTGAGGTCACATGCGTGGTGT GGACGTAAGCCACGAAGACCCTGAGGTCAAGTTCAACTGGTACGTGGACG GCGTGGAGGTGCATAATGCCAAGACAAAGCCGCGGGAGGAGCAGTACAAC 25 AGCACGTACCGTGTGGTCAGCGTCCTCACCGTCCTGCACCAGGACTGGCTG AATGGCAAGGAGTACAAGTGCAAGGTCTCCAACAAGCCCTCCCAACCCCC ATCGAGAAAACCATCTCCAAAGCCAAAGGGCAGCCCCGAGAACCACAGGT GTACACCCTGCCCCATCCCGGGATGAGCTGACCAAGAACCAGGTCAGCCT 30 GACCTGCCTGGTCAAAGGCTTCTATCCAAGCGACATCGCCGTGGAGTGGGA GAGCAATGGGCAGCCGGAGAACAACTACAAGACCACGCCTCCCGTGCTGG ACTCCGACGCTCCTTCTTCCTCTACAGCAAGCTCACCGTGGACAAGAGCA GGTGGCAGCAGGGGAACGTCTTCTCATGCTCCGTGATGCATGAGGCTCTGC ACAACCACTACACGCAGAAGAGCCTCTCCCTGTCTCCGGGTAAATGAGTGC 35 GACGGCCGCGACTCTAGAGGAT (SEQ ID NO:1)

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Example 10: Production of an Antibody from a Polypeptide

The antibodies of the present invention can be prepared by a variety of methods. (See, Current Protocols, Chapter 2.) For example, cells expressing a polypeptide of the present invention is administered to an animal to induce the production of sera containing polyclonal antibodies. In a preferred method, a preparation of the secreted protein is prepared and purified to render it substantially free of natural contaminants. Such a preparation is then introduced into an animal in order to produce polyclonal antisera of greater specific activity.

In the most preferred method, the antibodies of the present invention are monoclonal antibodies (or protein binding fragments thereof). Such monoclonal antibodies can be prepared using hybridoma technology. (Köhler et al., Nature 256:495 (1975); Köhler et al., Eur. J. Immunol. 6:511 (1976); Köhler et al., Eur. J. Immunol. 6:292 (1976); Hammerling et al., in: Monoclonal Antibodies and T-Cell Hybridomas, Elsevier, N.Y., pp. 563-681 (1981).) In general, such procedures involve immunizing an animal (preferably a mouse) with polypeptide or, more preferably, with a secreted polypeptide-expressing cell. Such cells may be cultured in any suitable tissue culture medium; however, it is preferable to culture cells in Earle's modified Eagle's medium supplemented with 10% fetal bovine serum (inactivated at about 56°C), and supplemented with about 10 g/l of nonessential amino acids, about 1,000 U/ml of penicillin, and about 100 μg/ml of streptomycin.

The splenocytes of such mice are extracted and fused with a suitable myeloma cell line. Any suitable myeloma cell line may be employed in accordance with the present invention; however, it is preferable to employ the parent myeloma cell line (SP2O), available from the ATCC. After fusion, the resulting hybridoma cells are selectively maintained in HAT medium, and then cloned by limiting dilution as described by Wands et al. (Gastroenterology 80:225-232 (1981).) The hybridoma cells obtained through such a selection are then assayed to identify clones which secrete antibodies capable of binding the polypeptide.

Alternatively, additional antibodies capable of binding to the polypeptide can be produced in a two-step procedure using anti-idiotypic antibodies. Such a method makes use of the fact that antibodies are themselves antigens, and therefore, it is possible to obtain an antibody which binds to a second antibody. In accordance with this method, protein specific antibodies are used to immunize an animal, preferably a mouse. The splenocytes of such an animal are then used to produce hybridoma cells, and the hybridoma cells are screened to identify clones which produce an antibody whose ability to bind to the protein-specific antibody can be blocked by the polypeptide.

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Such antibodies comprise anti-idiotypic antibodies to the protein-specific antibody and can be used to immunize an animal to induce formation of further protein-specific antibodies.

It will be appreciated that Fab and F(ab')2 and other fragments of the antibodies of the present invention may be used according to the methods disclosed herein. Such fragments are typically produced by proteolytic cleavage, using enzymes such as papain (to produce Fab fragments) or pepsin (to produce F(ab')2 fragments). Alternatively, secreted protein-binding fragments can be produced through the application of recombinant DNA technology or through synthetic chemistry.

For in vivo use of antibodies in humans, it may be preferable to use "humanized" chimeric monoclonal antibodies. Such antibodies can be produced using genetic constructs derived from hybridoma cells producing the monoclonal antibodies described above. Methods for producing chimeric antibodies are known in the art. (See, for review, Morrison, Science 229:1202 (1985); Oi et al., BioTechniques 4:214 (1986); Cabilly et al., U.S. Patent No. 4,816,567; Taniguchi et al., EP 171496; Morrison et al., EP 173494; Neuberger et al., WO 8601533; Robinson et al., WO 8702671; Boulianne et al., Nature 312:643 (1984); Neuberger et al., Nature 314:268 (1985).)

20 <u>Example 11: Production Of Secreted Protein For High-Throughput Screening Assays</u>

The following protocol produces a supernatant containing a polypeptide to be tested. This supernatant can then be used in the Screening Assays described in Examples 13-20.

First, dilute Poly-D-Lysine (644 587 Boehringer-Mannheim) stock solution (1mg/ml in PBS) 1:20 in PBS (w/o calcium or magnesium 17-516F Biowhittaker) for a working solution of 50ug/ml. Add 200 ul of this solution to each well (24 well plates) and incubate at RT for 20 minutes. Be sure to distribute the solution over each well (note: a 12-channel pipetter may be used with tips on every other channel). Aspirate off the Poly-D-Lysine solution and rinse with 1ml PBS (Phosphate Buffered Saline). The PBS should remain in the well until just prior to plating the cells and plates may be poly-lysine coated in advance for up to two weeks.

Plate 293T cells (do not carry cells past P+20) at 2 x 10⁵ cells/well in .5ml DMEM(Dulbecco's Modified Eagle Medium)(with 4.5 G/L glucose and L-glutamine (12-604F Biowhittaker))/10% heat inactivated FBS(14-503F Biowhittaker)/1x Penstrep(17-602E Biowhittaker). Let the cells grow overnight.

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The next day, mix together in a sterile solution basin: 300 ul Lipofectamine (18324-012 Gibco/BRL) and 5ml Optimem I (31985070 Gibco/BRL)/96-well plate. With a small volume multi-channel pipetter, aliquot approximately 2ug of an expression vector containing a polynucleotide insert, produced by the methods described in Examples 8 or 9, into an appropriately labeled 96-well round bottom plate. With a multi-channel pipetter, add 50ul of the Lipofectamine/Optimem I mixture to each well. Pipette up and down gently to mix. Incubate at RT 15-45 minutes. After about 20 minutes, use a multi-channel pipetter to add 150ul Optimem I to each well. As a control, one plate of vector DNA lacking an insert should be transfected with each set of transfections.

Preferably, the transfection should be performed by tag-teaming the following tasks. By tag-teaming, hands on time is cut in half, and the cells do not spend too much time on PBS. First, person A aspirates off the media from four 24-well plates of cells, and then person B rinses each well with .5-1ml PBS. Person A then aspirates off PBS rinse, and person B, using a12-channel pipetter with tips on every other channel, adds the 200ul of DNA/Lipofectamine/Optimem I complex to the odd wells first, then to the even wells, to each row on the 24-well plates. Incubate at 37°C for 6 hours.

While cells are incubating, prepare appropriate media, either 1%BSA in DMEM with 1x penstrep, or CHO-5 media (116.6 mg/L of CaCl2 (anhyd); 0.00130 mg/L $CuSO_4$ -5 H_2O ; 0.050 mg/L of $Fe(NO_3)_3$ -9 H_2O ; 0.417 mg/L of $FeSO_4$ -7 H_2O ; 311.80 20 mg/L of Kcl; 28.64 mg/L of MgCl₂; 48.84 mg/L of MgSO₄; 6995.50 mg/L of NaCl; 2400.0 mg/L of NaHCO₃; 62.50 mg/L of NaH₂PO₄-H₂0; 71.02 mg/L of Na₂HPO₄; .4320 mg/L of ZnSO₄-7H₂O; .002 mg/L of Arachidonic Acid; 1.022 mg/L of Cholesterol; .070 mg/L of DL-alpha-Tocopherol-Acetate; 0.0520 mg/L of Linoleic Acid: 0.010 mg/L of Linolenic Acid; 0.010 mg/L of Myristic Acid; 0.010 mg/L of Oleic 25 Acid; 0.010 mg/L of Palmitric Acid; 0.010 mg/L of Palmitic Acid; 100 mg/L of Pluronic F-68; 0.010 mg/L of Stearic Acid; 2.20 mg/L of Tween 80; 4551 mg/L of D-Glucose; 130.85 mg/ml of L- Alanine; 147.50 mg/ml of L-Arginine-HCL; 7.50 mg/ml of L-Asparagine-H,0; 6.65 mg/ml of L-Aspartic Acid; 29.56 mg/ml of L-Cystine-2HCL-H₂0; 31.29 mg/ml of L-Cystine-2HCL; 7.35 mg/ml of L-Glutamic Acid; 365.0 30 mg/ml of L-Glutamine; 18.75 mg/ml of Glycine; 52.48 mg/ml of L-Histidine-HCL-H₂0; 106.97 mg/ml of L-Isoleucine; 111.45 mg/ml of L-Leucine; 163.75 mg/ml of L-Lysine HCL; 32.34 mg/ml of L-Methionine; 68.48 mg/ml of L-Phenylalainine; 40.0 mg/ml of L-Proline; 26.25 mg/ml of L-Serine; 101.05 mg/ml of L-Threonine; 19.22 mg/ml of L-Tryptophan; 91.79 mg/ml of L-Tryrosine-2Na-2H,0; 99.65 mg/ml of L-35 Valine; 0.0035 mg/L of Biotin; 3.24 mg/L of D-Ca Pantothenate; 11.78 mg/L of

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Choline Chloride: 4.65 mg/L of Folic Acid; 15.60 mg/L of i-Inositol: 3.02 mg/L of Niacinamide; 3.00 mg/L of Pyridoxal HCL; 0.031 mg/L of Pyridoxine HCL: 0.319 mg/L of Riboflavin; 3.17 mg/L of Thiamine HCL; 0.365 mg/L of Thymidine; and 0.680 mg/L of Vitamin B₁₂; 25 mM of HEPES Buffer; 2.39 mg/L of Na Hypoxanthine; 0.105 mg/L of Lipoic Acid; 0.081 mg/L of Sodium Putrescine-2HCL; 55.0 mg/L of Sodium Pyruvate; 0.0067 mg/L of Sodium Selenite; 20uM of Ethanolamine; 0.122 mg/L of Ferric Citrate; 41.70 mg/L of Methyl-B-Cyclodextrin complexed with Linoleic Acid; 33.33 mg/L of Methyl-B-Cyclodextrin complexed with Oleic Acid; and 10 mg/L of Methyl-B-Cyclodextrin complexed with Retinal) with 2mm glutamine and 1x penstrep. (BSA (81-068-3 Bayer) 100gm dissolved in 1L DMEM for a 10% BSA stock solution). Filter the media and collect 50 ul for endotoxin assay in 15ml polystyrene conical.

The transfection reaction is terminated, preferably by tag-teaming, at the end of the incubation period. Person A aspirates off the transfection media, while person B adds 1.5ml appropriate media to each well. Incubate at 37°C for 45 or 72 hours depending on the media used: 1%BSA for 45 hours or CHO-5 for 72 hours.

On day four, using a 300ul multichannel pipetter, aliquot 600ul in one 1ml deep well plate and the remaining supernatant into a 2ml deep well. The supernatants from each well can then be used in the assays described in Examples 13-20.

It is specifically understood that when activity is obtained in any of the assays described below using a supernatant, the activity originates from either the polypeptide directly (e.g., as a secreted protein) or by the polypeptide inducing expression of other proteins, which are then secreted into the supernatant. Thus, the invention further provides a method of identifying the protein in the supernatant characterized by an activity in a particular assay.

Example 12: Construction of GAS Reporter Construct

One signal transduction pathway involved in the differentiation and proliferation of cells is called the Jaks-STATs pathway. Activated proteins in the Jaks-STATs pathway bind to gamma activation site "GAS" elements or interferon-sensitive responsive element ("ISRE"), located in the promoter of many genes. The binding of a protein to these elements alter the expression of the associated gene.

GAS and ISRE elements are recognized by a class of transcription factors called Signal Transducers and Activators of Transcription, or "STATs." There are six members of the STATs family. Stat1 and Stat3 are present in many cell types, as is Stat2 (as response to IFN-alpha is widespread). Stat4 is more restricted and is not in

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many cell types though it has been found in T helper class I, cells after treatment with IL-12. Stat5 was originally called mammary growth factor, but has been found at higher concentrations in other cells including myeloid cells. It can be activated in tissue culture cells by many cytokines.

The STATs are activated to translocate from the cytoplasm to the nucleus upon tyrosine phosphorylation by a set of kinases known as the Janus Kinase ("Jaks") family. Jaks represent a distinct family of soluble tyrosine kinases and include Tyk2, Jak1, Jak2, and Jak3. These kinases display significant sequence similarity and are generally catalytically inactive in resting cells.

The Jaks are activated by a wide range of receptors summarized in the Table below. (Adapted from review by Schidler and Darnell, Ann. Rev. Biochem. 64:621-51 (1995).) A cytokine receptor family, capable of activating Jaks, is divided into two groups: (a) Class 1 includes receptors for IL-2, IL-3, IL-4, IL-6, IL-7, IL-9, IL-11, IL-12, IL-15, Epo, PRL, GH, G-CSF, GM-CSF, LIF, CNTF, and thrombopoietin; and (b) Class 2 includes IFN-a, IFN-g, and IL-10. The Class 1 receptors share a conserved cysteine motif (a set of four conserved cysteines and one tryptophan) and a WSXWS motif (a membrane proxial region encoding Trp-Ser-Xxx-Trp-Ser (SEQ ID NO:2)).

Thus, on binding of a ligand to a receptor, Jaks are activated, which in turn activate STATs, which then translocate and bind to GAS elements. This entire process is encompassed in the Jaks-STATs signal transduction pathway.

Therefore, activation of the Jaks-STATs pathway, reflected by the binding of the GAS or the ISRE element, can be used to indicate proteins involved in the proliferation and differentiation of cells. For example, growth factors and cytokines are known to activate the Jaks-STATs pathway. (See Table below.) Thus, by using GAS elements linked to reporter molecules, activators of the Jaks-STATs pathway can be identified.

	<u>Ligand</u>	tyk2	JAKs Jak l	<u>Jak2</u>	Jak3	<u>STATS</u>	GAS(elements) or ISRE
5	IFN family IFN-a/B IFN-g Il-10	+	+ + ?	- + ?	- - -	1,2,3 1 1,3	ISRE GAS (IRF1>Lys6>IFP)
10	gp130 family IL-6 (Pleiotrohic) Il-11(Pleiotrohic) OnM(Pleiotrohic)	+ ? ?	+ + +	+ ? +	????	1,3 1,3 1,3	GAS (IRF1>Lys6>IFP)
15	LIF(Pleiotrohic) CNTF(Pleiotrohic) G-CSF(Pleiotrohic) IL-12(Pleiotrohic)	? -/+ ? +	+ + + -	+ + ? +	? ? ? +	1,3 1,3 1,3 1,3	
20	g-C family IL-2 (lymphocytes) IL-4 (lymph/myeloid) IL-7 (lymphocytes) IL-9 (lymphocytes) IL-13 (lymphocyte) IL-15	- - - - -	+ + + + +	- - - ?	+ + + + ?	1,3,5 6 5 5 6 5	GAS GAS (IRF1 = IFP >>Ly6)(IgH) GAS GAS GAS GAS GAS
25 30	gp140 family IL-3 (myeloid) IL-5 (myeloid) GM-CSF (myeloid)	- - -	- - -	+ + +	- -	5 5 5 5	GAS (IRF1>IFP>>Ly6) GAS GAS
35	Growth hormone fami GH PRL EPO	il <u>y</u> ? ? ?	- +/- -	+ + +	-	5 1,3,5 5	GAS(B-CAS>IRF1=IFP>>Ly6)
40	Receptor Tyrosine Kin EGF PDGF CSF-1	nases ? ? ?	++++++	+ + +	-	1,3 1,3 1,3	GAS (IRF1) GAS (not IRF1)

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To construct a synthetic GAS containing promoter element, which is used in the Biological Assays described in Examples 13-14, a PCR based strategy is employed to generate a GAS-SV40 promoter sequence. The 5' primer contains four tandem copies of the GAS binding site found in the IRF1 promoter and previously demonstrated to bind STATs upon induction with a range of cytokines (Rothman et al., Immunity 1:457-468 (1994).), although other GAS or ISRE elements can be used instead. The 5' primer also contains 18bp of sequence complementary to the SV40 early promoter sequence and is flanked with an XhoI site. The sequence of the 5' primer is: 5':GCGCCTCGAGATTTCCCCGAAATCTAGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTC

The downstream primer is complementary to the SV40 promoter and is flanked with a Hind III site: 5':GCGGCAAGCTTTTTGCAAAGCCTAGGC:3' (SEQ ID NO:4)

PCR amplification is performed using the SV40 promoter template present in the B-gal:promoter plasmid obtained from Clontech. The resulting PCR fragment is digested with XhoI/Hind III and subcloned into BLSK2-. (Stratagene.) Sequencing with forward and reverse primers confirms that the insert contains the following sequence:

5':CTCGAGATTTCCCCGAAATCTAGATTTCCCCGAAATGATTTCCCCGAAATG
ATTTCCCCGAAATATCTGCCATCTCAATTAGTCAGCAACCATAGTCCCGCCC
CTAACTCCGCCCATCCCGCCCCTAACTCCGCCCAGTTCCGCCCATTCTCCGC
CCCATGGCTGACTAATTTTTTTTATTTATTTATGCAGAGGCCGAGGCCGCCTCGGC
CTCTGAGCTATTCCAGAAGTAGTGAGGAGGCTTTTTTTGGAGGCCTAGGCTTT
TGCAAAAAAGCTT:3' (SEQ ID NO:5)

With this GAS promoter element linked to the SV40 promoter, a GAS:SEAP2 reporter construct is next engineered. Here, the reporter molecule is a secreted alkaline phosphatase, or "SEAP." Clearly, however, any reporter molecule can be instead of SEAP, in this or in any of the other Examples. Well known reporter molecules that can be used instead of SEAP include chloramphenicol acetyltransferase (CAT), luciferase, alkaline phosphatase, B-galactosidase, green fluorescent protein (GFP), or any protein detectable by an antibody.

The above sequence confirmed synthetic GAS-SV40 promoter element is subcloned into the pSEAP-Promoter vector obtained from Clontech using HindIII and XhoI, effectively replacing the SV40 promoter with the amplified GAS:SV40 promoter element, to create the GAS-SEAP vector. However, this vector does not contain a neomycin resistance gene, and therefore, is not preferred for mammalian expression systems.

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Thus, in order to generate mammalian stable cell lines expressing the GAS-SEAP reporter, the GAS-SEAP cassette is removed from the GAS-SEAP vector using SalI and NotI, and inserted into a backbone vector containing the neomycin resistance gene, such as pGFP-1 (Clontech), using these restriction sites in the multiple cloning site, to create the GAS-SEAP/Neo vector. Once this vector is transfected into mammalian cells, this vector can then be used as a reporter molecule for GAS binding as described in Examples 13-14.

Other constructs can be made using the above description and replacing GAS with a different promoter sequence. For example, construction of reporter molecules containing NFK-B and EGR promoter sequences are described in Examples 15 and 16. However, many other promoters can be substituted using the protocols described in these Examples. For instance, SRE, IL-2, NFAT, or Osteocalcin promoters can be substituted, alone or in combination (e.g., GAS/NF-KB/EGR, GAS/NF-KB, Il-2/NFAT, or NF-KB/GAS). Similarly, other cell lines can be used to test reporter construct activity, such as HELA (epithelial), HUVEC (endothelial), Reh (B-cell), Saos-2 (osteoblast), HUVAC (aortic), or Cardiomyocyte.

Example 13: High-Throughput Screening Assay for T-cell Activity.

The following protocol is used to assess T-cell activity by identifying factors, such as growth factors and cytokines, that may proliferate or differentiate T-cells. T-cell activity is assessed using the GAS/SEAP/Neo construct produced in Example 12. Thus, factors that increase SEAP activity indicate the ability to activate the Jaks-STATS signal transduction pathway. The T-cell used in this assay is Jurkat T-cells (ATCC Accession No. TIB-152), although Molt-3 cells (ATCC Accession No. CRL-1552) and Molt-4 cells (ATCC Accession No. CRL-1582) cells can also be used.

Jurkat T-cells are lymphoblastic CD4+ Th1 helper cells. In order to generate stable cell lines, approximately 2 million Jurkat cells are transfected with the GAS-SEAP/neo vector using DMRIE-C (Life Technologies)(transfection procedure described below). The transfected cells are seeded to a density of approximately 20,000 cells per well and transfectants resistant to 1 mg/ml genticin selected. Resistant colonies are expanded and then tested for their response to increasing concentrations of interferon gamma. The dose response of a selected clone is demonstrated.

Specifically, the following protocol will yield sufficient cells for 75 wells containing 200 ul of cells. Thus, it is either scaled up, or performed in multiple to generate sufficient cells for multiple 96 well plates. Jurkat cells are maintained in RPMI + 10% serum with 1%Pen-Strep. Combine 2.5 mls of OPTI-MEM (Life Technologies)

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with 10 ug of plasmid DNA in a T25 flask. Add 2.5 ml OPTI-MEM containing 50 ul of DMRIE-C and incubate at room temperature for 15-45 mins.

During the incubation period, count cell concentration, spin down the required number of cells (10^7 per transfection), and resuspend in OPTI-MEM to a final concentration of 10^7 cells/ml. Then add 1ml of 1 x 10^7 cells in OPTI-MEM to T25 flask and incubate at 37°C for 6 hrs. After the incubation, add 10 ml of RPMI + 15% serum.

The Jurkat:GAS-SEAP stable reporter lines are maintained in RPMI + 10% serum, 1 mg/ml Genticin, and 1% Pen-Strep. These cells are treated with supernatants containing a polypeptide as produced by the protocol described in Example 11.

On the day of treatment with the supernatant, the cells should be washed and resuspended in fresh RPMI + 10% serum to a density of 500,000 cells per ml. The exact number of cells required will depend on the number of supernatants being screened. For one 96 well plate, approximately 10 million cells (for 10 plates, 100 million cells) are required.

Transfer the cells to a triangular reservoir boat, in order to dispense the cells into a 96 well dish, using a 12 channel pipette. Using a 12 channel pipette, transfer 200 ul of cells into each well (therefore adding 100, 000 cells per well).

After all the plates have been seeded, 50 ul of the supernatants are transferred directly from the 96 well plate containing the supernatants into each well using a 12 channel pipette. In addition, a dose of exogenous interferon gamma (0.1, 1.0, 10 ng) is added to wells H9, H10, and H11 to serve as additional positive controls for the assay.

The 96 well dishes containing Jurkat cells treated with supernatants are placed in an incubator for 48 hrs (note: this time is variable between 48-72 hrs). 35 ul samples from each well are then transferred to an opaque 96 well plate using a 12 channel pipette. The opaque plates should be covered (using sellophene covers) and stored at -20°C until SEAP assays are performed according to Example 17. The plates containing the remaining treated cells are placed at 4°C and serve as a source of material for repeating the assay on a specific well if desired.

As a positive control, 100 Unit/ml interferon gamma can be used which is known to activate Jurkat T cells. Over 30 fold induction is typically observed in the positive control wells.

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Example 14: High-Throughput Screening Assay Identifying Myeloid Activity

The following protocol is used to assess myeloid activity by identifying factors, such as growth factors and cytokines, that may proliferate or differentiate myeloid cells. Myeloid cell activity is assessed using the GAS/SEAP/Neo construct produced in Example 12. Thus, factors that increase SEAP activity indicate the ability to activate the Jaks-STATS signal transduction pathway. The myeloid cell used in this assay is U937, a pre-monocyte cell line, although TF-1, HL60, or KG1 can be used.

To transiently transfect U937 cells with the GAS/SEAP/Neo construct produced in Example 12, a DEAE-Dextran method (Kharbanda et. al., 1994, Cell Growth & Differentiation, 5:259-265) is used. First, harvest 2x10e⁷ U937 cells and wash with PBS. The U937 cells are usually grown in RPMI 1640 medium containing 10% heat-inactivated fetal bovine serum (FBS) supplemented with 100 units/ml penicillin and 100 mg/ml streptomycin.

Next, suspend the cells in 1 ml of 20 mM Tris-HCl (pH 7.4) buffer containing 0.5 mg/ml DEAE-Dextran, 8 ug GAS-SEAP2 plasmid DNA, 140 mM NaCl, 5 mM KCl, 375 uM Na₂HPO₄.7H₂O, 1 mM MgCl₂, and 675 uM CaCl₂. Incubate at 37°C for 45 min.

Wash the cells with RPMI 1640 medium containing 10% FBS and then resuspend in 10 ml complete medium and incubate at 37°C for 36 hr.

The GAS-SEAP/U937 stable cells are obtained by growing the cells in 400 ug/ml G418. The G418-free medium is used for routine growth but every one to two months, the cells should be re-grown in 400 ug/ml G418 for couple of passages.

These cells are tested by harvesting $1x10^8$ cells (this is enough for ten 96-well plates assay) and wash with PBS. Suspend the cells in 200 ml above described growth medium, with a final density of $5x10^5$ cells/ml. Plate 200 ul cells per well in the 96-well plate (or $1x10^5$ cells/well).

Add 50 ul of the supernatant prepared by the protocol described in Example 11. Incubate at 37°C for 48 to 72 hr. As a positive control, 100 Unit/ml interferon gamma can be used which is known to activate U937 cells. Over 30 fold induction is typically observed in the positive control wells. SEAP assay the supernatant according to the protocol described in Example 17.

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Example 15: High-Throughput Screening Assay Identifying Neuronal Activity.

When cells undergo differentiation and proliferation, a group of genes are activated through many different signal transduction pathways. One of these genes, EGR1 (early growth response gene 1), is induced in various tissues and cell types upon activation. The promoter of EGR1 is responsible for such induction. Using the EGR1 promoter linked to reporter molecules, activation of cells can be assessed.

Particularly, the following protocol is used to assess neuronal activity in PC12 cell lines. PC12 cells (rat phenochromocytoma cells) are known to proliferate and/or differentiate by activation with a number of mitogens, such as TPA (tetradecanoyl phorbol acetate), NGF (nerve growth factor), and EGF (epidermal growth factor). The EGR1 gene expression is activated during this treatment. Thus, by stably transfecting PC12 cells with a construct containing an EGR promoter linked to SEAP reporter, activation of PC12 cells can be assessed.

The EGR/SEAP reporter construct can be assembled by the following protocol. The EGR-1 promoter sequence (-633 to +1)(Sakamoto K et al., Oncogene 6:867-871 (1991)) can be PCR amplified from human genomic DNA using the following primers:

- 5' GCGCTCGAGGGATGACAGCGATAGAACCCCGG -3' (SEQ ID NO:6)
- 5' GCGAAGCTTCGCGACTCCCGGATCCGCCTC-3' (SEQ ID NO:7)

Using the GAS:SEAP/Neo vector produced in Example 12, EGR1 amplified product can then be inserted into this vector. Linearize the GAS:SEAP/Neo vector using restriction enzymes XhoI/HindIII, removing the GAS/SV40 stuffer. Restrict the EGR1 amplified product with these same enzymes. Ligate the vector and the EGR1 promoter.

To prepare 96 well-plates for cell culture, two mls of a coating solution (1:30 dilution of collagen type I (Upstate Biotech Inc. Cat#08-115) in 30% ethanol (filter sterilized)) is added per one 10 cm plate or 50 ml per well of the 96-well plate, and allowed to air dry for 2 hr.

PC12 cells are routinely grown in RPMI-1640 medium (Bio Whittaker) containing 10% horse serum (JRH BIOSCIENCES, Cat. # 12449-78P), 5% heatinactivated fetal bovine serum (FBS) supplemented with 100 units/ml penicillin and 100 ug/ml streptomycin on a precoated 10 cm tissue culture dish. One to four split is done every three to four days. Cells are removed from the plates by scraping and resuspended with pipetting up and down for more than 15 times.

Transfect the EGR/SEAP/Neo construct into PC12 using the Lipofectamine protocol described in Example 11. EGR-SEAP/PC12 stable cells are obtained by growing the cells in 300 ug/ml G418. The G418-free medium is used for routine

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growth but every one to two months, the cells should be re-grown in 300 ug/ml G418 for couple of passages.

To assay for neuronal activity, a 10 cm plate with cells around 70 to 80% confluent is screened by removing the old medium. Wash the cells once with PBS (Phosphate buffered saline). Then starve the cells in low serum medium (RPMI-1640 containing 1% horse serum and 0.5% FBS with antibiotics) overnight.

The next morning, remove the medium and wash the cells with PBS. Scrape off the cells from the plate, suspend the cells well in 2 ml low serum medium. Count the cell number and add more low serum medium to reach final cell density as $5x10^5$ cells/ml.

Add 200 ul of the cell suspension to each well of 96-well plate (equivalent to $1x10^5$ cells/well). Add 50 ul supernatant produced by Example 11, 37°C for 48 to 72 hr. As a positive control, a growth factor known to activate PC12 cells through EGR can be used, such as 50 ng/ul of Neuronal Growth Factor (NGF). Over fifty-fold induction of SEAP is typically seen in the positive control wells. SEAP assay the supernatant according to Example 17.

Example 16: High-Throughput Screening Assay for T-cell Activity

NF-κB (Nuclear Factor κB) is a transcription factor activated by a wide variety of agents including the inflammatory cytokines IL-1 and TNF, CD30 and CD40, lymphotoxin-alpha and lymphotoxin-beta, by exposure to LPS or thrombin, and by expression of certain viral gene products. As a transcription factor, NF-κB regulates the expression of genes involved in immune cell activation, control of apoptosis (NF-κB appears to shield cells from apoptosis), B and T-cell development, anti-viral and antimicrobial responses, and multiple stress responses.

In non-stimulated conditions, NF- κB is retained in the cytoplasm with I- κB (Inhibitor κB). However, upon stimulation, I- κB is phosphorylated and degraded, causing NF- κB to shuttle to the nucleus, thereby activating transcription of target genes. Target genes activated by NF- κB include IL-2, IL-6, GM-CSF, ICAM-1 and class 1 MHC.

Due to its central role and ability to respond to a range of stimuli, reporter constructs utilizing the NF-kB promoter element are used to screen the supernatants produced in Example 11. Activators or inhibitors of NF-kB would be useful in treating

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diseases. For example, inhibitors of NF-kB could be used to treat those diseases related to the acute or chronic activation of NF-kB, such as rheumatoid arthritis.

To construct a vector containing the NF-κB promoter element, a PCR based strategy is employed. The upstream primer contains four tandem copies of the NF-κB binding site (GGGGACTTTCCC) (SEQ ID NO:8), 18 bp of sequence complementary to the 5' end of the SV40 early promoter sequence, and is flanked with an XhoI site: 5':GCGGCCTCGAGGGGACTTTCCCGGGGACTTTCCGGGACTTTCCGGGACTTTCCGGGACTTTCCGGGACTTTCCGGGACTTTCCGGAGACTTTCCGGGACTTTCCGGGACTTTCCGGGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGAGACTTTCGAGAACTTTCAACTTCAACTTCAACTTCAACTTCAACTTTCAACTTCAACTTCAACTTCAACTTCAACTTCAACTTCAACTTCAACTTCAACTTCAAC

The downstream primer is complementary to the 3' end of the SV40 promoter and is flanked with a Hind III site:

5':GCGGCAAGCTTTTTGCAAAGCCTAGGC:3' (SEQ ID NO:4)

PCR amplification is performed using the SV40 promoter template present in the pB-gal:promoter plasmid obtained from Clontech. The resulting PCR fragment is digested with XhoI and Hind III and subcloned into BLSK2-. (Stratagene) Sequencing with the T7 and T3 primers confirms the insert contains the following sequence:

5':CTCGAGGGGACTTTCCCGGGGACTTTCCGGGGACTTTCC ATCTGCCATCTCAATTAGTCAGCAACCATAGTCCCGCCCCTAACTCCGCCCA TCCCGCCCCTAACTCCGCCCAGTTCCGCCCATTCTCCGCCCCATGGCTGACT AATTTTTTTATTTATGCAGAGGCCGAGGCCGCCTCGGCCTCTGAGCTATTC CAGAAGTAGTGAGGAGGCTTTTTTGGAGGCCTAGGCTTTTGCAAAAAGCTT: 3' (SEQ ID NO:10)

Next, replace the SV40 minimal promoter element present in the pSEAP2-promoter plasmid (Clontech) with this NF-kB/SV40 fragment using XhoI and HindIII. However, this vector does not contain a neomycin resistance gene, and therefore, is not preferred for mammalian expression systems.

In order to generate stable mammalian cell lines, the NF-κB/SV40/SEAP

cassette is removed from the above NF-κB/SEAP vector using restriction enzymes SalI and NotI, and inserted into a vector containing neomycin resistance. Particularly, the NF-κB/SV40/SEAP cassette was inserted into pGFP-1 (Clontech), replacing the GFP gene, after restricting pGFP-1 with SalI and NotI.

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Once NF-kB/SV40/SEAP/Neo vector is created, stable Jurkat T-cells are created and maintained according to the protocol described in Example 13. Similarly, the method for assaying supernatants with these stable Jurkat T-cells is also described in Example 13. As a positive control, exogenous TNF alpha (0.1,1, 10 ng) is added to wells H9, H10, and H11, with a 5-10 fold activation typically observed.

Example 17: Assay for SEAP Activity

As a reporter molecule for the assays described in Examples 13-16, SEAP activity is assayed using the Tropix Phospho-light Kit (Cat. BP-400) according to the following general procedure. The Tropix Phospho-light Kit supplies the Dilution, Assay, and Reaction Buffers used below.

Prime a dispenser with the 2.5x Dilution Buffer and dispense 15 μ l of 2.5x dilution buffer into Optiplates containing 35 μ l of a supernatant. Seal the plates with a plastic sealer and incubate at 65°C for 30 min. Separate the Optiplates to avoid uneven heating.

Cool the samples to room temperature for 15 minutes. Empty the dispenser and prime with the Assay Buffer. Add 50 µl Assay Buffer and incubate at room temperature 5 min. Empty the dispenser and prime with the Reaction Buffer (see the table below). Add 50 µl Reaction Buffer and incubate at room temperature for 20 minutes. Since the intensity of the chemiluminescent signal is time dependent, and it takes about 10 minutes to read 5 plates on luminometer, one should treat 5 plates at each time and start the second set 10 minutes later.

Read the relative light unit in the luminometer. Set H12 as blank, and print the results. An increase in chemiluminescence indicates reporter activity.

Reaction Buffer Formulation:

Reaction B	utter Formulation:		
# of plates	Rxn buffer diluent (ml)	CSPD (ml)	_
10	60	3	_
11	65	3.25	
12	70	3.5	
13	75	3.75	
14	80	4	
15	85	4.25	
16	90	4.5	
17	95	4.75	
18	100	5	
19	105	5.25	
20	110	5.5	
21	115	5.75	
22	120	6	

23	125	6.25
24	130	6.5
25	135	6.75
26	140	7
27	145	7.25
28	150	7.5
29	155	7.75
30	160	8
31	165	8.25
32	170	8.5
33	175	8.75
34	180	9
35	185	9.25
36	190	9.5
37	195	9.75
38	200	10
39	205	10.25
40	210	10.5
41	215	10.75
42	220	11
43	225	11.25
44	230	11.5
45	235	11.75
46	240	12
47	245	12.25
48	250	12.5
49	255	12.75
50	260	13

Example 18: High-Throughput Screening Assay Identifying Changes in Small Molecule Concentration and Membrane Permeability

Binding of a ligand to a receptor is known to alter intracellular levels of small molecules, such as calcium, potassium, sodium, and pH, as well as alter membrane potential. These alterations can be measured in an assay to identify supernatants which bind to receptors of a particular cell. Although the following protocol describes an assay for calcium, this protocol can easily be modified to detect changes in potassium, sodium, pH, membrane potential, or any other small molecule which is detectable by a fluorescent probe.

The following assay uses Fluorometric Imaging Plate Reader ("FLIPR") to measure changes in fluorescent molecules (Molecular Probes) that bind small molecules. Clearly, any fluorescent molecule detecting a small molecule can be used instead of the calcium fluorescent molecule, fluo-3, used here.

For adherent cells, seed the cells at 10,000-20,000 cells/well in a Co-star black 96-well plate with clear bottom. The plate is incubated in a CO_2 incubator for 20 hours. The adherent cells are washed two times in Biotek washer with 200 ul of HBSS (Hank's Balanced Salt Solution) leaving 100 ul of buffer after the final wash.

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A stock solution of 1 mg/ml fluo-3 is made in 10% pluronic acid DMSO. To load the cells with fluo-3, 50 ul of 12 ug/ml fluo-3 is added to each well. The plate is incubated at 37° C in a CO_2 incubator for 60 min. The plate is washed four times in the Biotek washer with HBSS leaving 100 ul of buffer.

For non-adherent cells, the cells are spun down from culture media. Cells are re-suspended to $2-5\times10^6$ cells/ml with HBSS in a 50-ml conical tube. 4 ul of 1 mg/ml fluo-3 solution in 10% pluronic acid DMSO is added to each ml of cell suspension. The tube is then placed in a 37°C water bath for 30-60 min. The cells are washed twice with HBSS, resuspended to 1×10^6 cells/ml, and dispensed into a microplate, 100 ul/well. The plate is centrifuged at 1000 rpm for 5 min. The plate is then washed once in Denley CellWash with 200 ul, followed by an aspiration step to 100 ul final volume.

For a non-cell based assay, each well contains a fluorescent molecule, such as fluo-3. The supernatant is added to the well, and a change in fluorescence is detected.

To measure the fluorescence of intracellular calcium, the FLIPR is set for the following parameters: (1) System gain is 300-800 mW; (2) Exposure time is 0.4 second; (3) Camera F/stop is F/2; (4) Excitation is 488 nm; (5) Emission is 530 nm; and (6) Sample addition is 50 ul. Increased emission at 530 nm indicates an extracellular signaling event which has resulted in an increase in the intracellular Ca⁺⁺ concentration.

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Example 19: High-Throughput Screening Assay Identifying Tyrosine Kinase Activity

The Protein Tyrosine Kinases (PTK) represent a diverse group of transmembrane and cytoplasmic kinases. Within the Receptor Protein Tyrosine Kinase RPTK) group are receptors for a range of mitogenic and metabolic growth factors including the PDGF, FGF, EGF, NGF, HGF and Insulin receptor subfamilies. In addition there are a large family of RPTKs for which the corresponding ligand is unknown. Ligands for RPTKs include mainly secreted small proteins, but also membrane-bound and extracellular matrix proteins.

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Activation of RPTK by ligands involves ligand-mediated receptor dimerization, resulting in transphosphorylation of the receptor subunits and activation of the cytoplasmic tyrosine kinases. The cytoplasmic tyrosine kinases include receptor associated tyrosine kinases of the src-family (e.g., src, yes, lck, lyn, fyn) and non-receptor linked and cytosolic protein tyrosine kinases, such as the Jak family, members of which mediate signal transduction triggered by the cytokine superfamily of receptors (e.g., the Interleukins, Interferons, GM-CSF, and Leptin).

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Because of the wide range of known factors capable of stimulating tyrosine kinase activity, the identification of novel human secreted proteins capable of activating tyrosine kinase signal transduction pathways are of interest. Therefore, the following protocol is designed to identify those novel human secreted proteins capable of activating the tyrosine kinase signal transduction pathways.

Seed target cells (e.g., primary keratinocytes) at a density of approximately 25,000 cells per well in a 96 well Loprodyne Silent Screen Plates purchased from Nalge Nunc (Naperville, IL). The plates are sterilized with two 30 minute rinses with 100% ethanol, rinsed with water and dried overnight. Some plates are coated for 2 hr with 100 ml of cell culture grade type I collagen (50 mg/ml), gelatin (2%) or polylysine (50 mg/ml), all of which can be purchased from Sigma Chemicals (St. Louis, MO) or 10% Matrigel purchased from Becton Dickinson (Bedford,MA), or calf serum, rinsed with PBS and stored at 4°C. Cell growth on these plates is assayed by seeding 5,000 cells/well in growth medium and indirect quantitation of cell number through use of alamarBlue as described by the manufacturer Alamar Biosciences, Inc. (Sacramento, CA) after 48 hr. Falcon plate covers #3071 from Becton Dickinson (Bedford,MA) are used to cover the Loprodyne Silent Screen Plates. Falcon Microtest III cell culture plates can also be used in some proliferation experiments.

To prepare extracts, A431 cells are seeded onto the nylon membranes of Loprodyne plates (20,000/200ml/well) and cultured overnight in complete medium. Cells are quiesced by incubation in serum-free basal medium for 24 hr. After 5-20 minutes treatment with EGF (60ng/ml) or 50 ul of the supernatant produced in Example 11, the medium was removed and 100 ml of extraction buffer ((20 mM HEPES pH 7.5, 0.15 M NaCl, 1% Triton X-100, 0.1% SDS, 2 mM Na3VO4, 2 mM Na4P2O7 and a cocktail of protease inhibitors (# 1836170) obtained from Boeheringer Mannheim (Indianapolis, IN) is added to each well and the plate is shaken on a rotating shaker for 5 minutes at 4°C. The plate is then placed in a vacuum transfer manifold and the extract filtered through the 0.45 mm membrane bottoms of each well using house vacuum. Extracts are collected in a 96-well catch/assay plate in the bottom of the vacuum manifold and immediately placed on ice. To obtain extracts clarified by centrifugation, the content of each well, after detergent solubilization for 5 minutes, is removed and centrifuged for 15 minutes at 4°C at 16,000 x g.

Test the filtered extracts for levels of tyrosine kinase activity. Although many methods of detecting tyrosine kinase activity are known, one method is described here.

Generally, the tyrosine kinase activity of a supernatant is evaluated by determining its ability to phosphorylate a tyrosine residue on a specific substrate (a

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biotinylated peptide). Biotinylated peptides that can be used for this purpose include PSK1 (corresponding to amino acids 6-20 of the cell division kinase cdc2-p34) and PSK2 (corresponding to amino acids 1-17 of gastrin). Both peptides are substrates for a range of tyrosine kinases and are available from Boehringer Mannheim.

The tyrosine kinase reaction is set up by adding the following components in order. First, add 10ul of 5uM Biotinylated Peptide, then 10ul ATP/Mg₂₊ (5mM ATP/50mM MgCl₂), then 10ul of 5x Assay Buffer (40mM imidazole hydrochloride, pH7.3, 40 mM beta-glycerophosphate, 1mM EGTA, 100mM MgCl₂, 5 mM MnCl₂, 0.5 mg/ml BSA), then 5ul of Sodium Vanadate(1mM), and then 5ul of water. Mix the components gently and preincubate the reaction mix at 30°C for 2 min. Initial the reaction by adding 10ul of the control enzyme or the filtered supernatant.

The tyrosine kinase assay reaction is then terminated by adding 10 ul of 120mm EDTA and place the reactions on ice.

Tyrosine kinase activity is determined by transferring 50 ul aliquot of reaction mixture to a microtiter plate (MTP) module and incubating at 37°C for 20 min. This allows the streptavadin coated 96 well plate to associate with the biotinylated peptide. Wash the MTP module with 300ul/well of PBS four times. Next add 75 ul of anti-phospotyrosine antibody conjugated to horse radish peroxidase(anti-P-Tyr-POD(0.5u/ml)) to each well and incubate at 37°C for one hour. Wash the well as above.

Next add 100ul of peroxidase substrate solution (Boehringer Mannheim) and incubate at room temperature for at least 5 mins (up to 30 min). Measure the absorbance of the sample at 405 nm by using ELISA reader. The level of bound peroxidase activity is quantitated using an ELISA reader and reflects the level of tyrosine kinase activity.

Example 20: High-Throughput Screening Assay Identifying Phosphorvlation Activity

As a potential alternative and/or compliment to the assay of protein tyrosine kinase activity described in Example 19, an assay which detects activation (phosphorylation) of major intracellular signal transduction intermediates can also be used. For example, as described below one particular assay can detect tyrosine phosphorylation of the Erk-1 and Erk-2 kinases. However, phosphorylation of other molecules, such as Raf, JNK, p38 MAP, Map kinase kinase (MEK), MEK kinase, Src, Muscle specific kinase (MuSK), IRAK, Tec, and Janus, as well as any other

phosphoserine, phosphotyrosine, or phosphothreonine molecule, can be detected by substituting these molecules for Erk-1 or Erk-2 in the following assay.

Specifically, assay plates are made by coating the wells of a 96-well ELISA plate with 0.1ml of protein G (lug/ml) for 2 hr at room temp, (RT). The plates are then rinsed with PBS and blocked with 3% BSA/PBS for 1 hr at RT. The protein G plates are then treated with 2 commercial monoclonal antibodies (100ng/well) against Erk-1 and Erk-2 (1 hr at RT) (Santa Cruz Biotechnology). (To detect other molecules, this step can easily be modified by substituting a monoclonal antibody detecting any of the above described molecules.) After 3-5 rinses with PBS, the plates are stored at 4°C until use.

A431 cells are seeded at 20,000/well in a 96-well Loprodyne filterplate and cultured overnight in growth medium. The cells are then starved for 48 hr in basal medium (DMEM) and then treated with EGF (6ng/well) or 50 ul of the supernatants obtained in Example 11 for 5-20 minutes. The cells are then solubilized and extracts filtered directly into the assay plate.

After incubation with the extract for 1 hr at RT, the wells are again rinsed. As a positive control, a commercial preparation of MAP kinase (10ng/well) is used in place of A431 extract. Plates are then treated with a commercial polyclonal (rabbit) antibody (1ug/ml) which specifically recognizes the phosphorylated epitope of the Erk-1 and Erk-2 kinases (1 hr at RT). This antibody is biotinylated by standard procedures. The bound polyclonal antibody is then quantitated by successive incubations with Europium-streptavidin and Europium fluorescence enhancing reagent in the Wallac DELFIA instrument (time-resolved fluorescence). An increased fluorescent signal over background indicates a phosphorylation.

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Example 21: Method of Determining Alterations in a Gene Corresponding to a Polynucleotide

RNA isolated from entire families or individual patients presenting with a phenotype of interest (such as a disease) is be isolated. cDNA is then generated from these RNA samples using protocols known in the art. (See, Sambrook.) The cDNA is then used as a template for PCR, employing primers surrounding regions of interest in SEQ ID NO:X. Suggested PCR conditions consist of 35 cycles at 95°C for 30 seconds; 60-120 seconds at 52-58°C; and 60-120 seconds at 70°C, using buffer solutions described in Sidransky, D., et al., Science 252:706 (1991).

PCR products are then sequenced using primers labeled at their 5' end with T4 polynucleotide kinase, employing SequiTherm Polymerase. (Epicentre Technologies).

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The intron-exon borders of selected exons is also determined and genomic PCR products analyzed to confirm the results. PCR products harboring suspected mutations is then cloned and sequenced to validate the results of the direct sequencing.

PCR products is cloned into T-tailed vectors as described in Holton, T.A. and Graham, M.W., Nucleic Acids Research, 19:1156 (1991) and sequenced with T7 polymerase (United States Biochemical). Affected individuals are identified by mutations not present in unaffected individuals.

Genomic rearrangements are also observed as a method of determining alterations in a gene corresponding to a polynucleotide. Genomic clones isolated according to Example 2 are nick-translated with digoxigenindeoxy-uridine 5'-triphosphate (Boehringer Manheim), and FISH performed as described in Johnson, Cg. et al., Methods Cell Biol. 35:73-99 (1991). Hybridization with the labeled probe is carried out using a vast excess of human cot-1 DNA for specific hybridization to the corresponding genomic locus.

Chromosomes are counterstained with 4,6-diamino-2-phenylidole and propidium iodide, producing a combination of C- and R-bands. Aligned images for precise mapping are obtained using a triple-band filter set (Chroma Technology, Brattleboro, VT) in combination with a cooled charge-coupled device camera (Photometrics, Tucson, AZ) and variable excitation wavelength filters. (Johnson, Cv. et al., Genet. Anal. Tech. Appl., 8:75 (1991).) Image collection, analysis and chromosomal fractional length measurements are performed using the ISee Graphical Program System. (Inovision Corporation, Durham, NC.) Chromosome alterations of the genomic region hybridized by the probe are identified as insertions, deletions, and translocations. These alterations are used as a diagnostic marker for an associated disease.

Example 22: Method of Detecting Abnormal Levels of a Polypeptide in a Biological Sample

A polypeptide of the present invention can be detected in a biological sample, and if an increased or decreased level of the polypeptide is detected, this polypeptide is a marker for a particular phenotype. Methods of detection are numerous, and thus, it is understood that one skilled in the art can modify the following assay to fit their particular needs.

For example, antibody-sandwich ELISAs are used to detect polypeptides in a sample, preferably a biological sample. Wells of a microtiter plate are coated with specific antibodies, at a final concentration of 0.2 to 10 ug/ml. The antibodies are either monoclonal or polyclonal and are produced by the method described in Example 10.

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The wells are blocked so that non-specific binding of the polypeptide to the well is reduced.

The coated wells are then incubated for > 2 hours at RT with a sample containing the polypeptide. Preferably, serial dilutions of the sample should be used to validate results. The plates are then washed three times with deionized or distilled water to remove unbounded polypeptide.

Next, 50 ul of specific antibody-alkaline phosphatase conjugate, at a concentration of 25-400 ng, is added and incubated for 2 hours at room temperature. The plates are again washed three times with deionized or distilled water to remove unbounded conjugate.

Add 75 ul of 4-methylumbelliferyl phosphate (MUP) or p-nitrophenyl phosphate (NPP) substrate solution to each well and incubate 1 hour at room temperature. Measure the reaction by a microtiter plate reader. Prepare a standard curve, using serial dilutions of a control sample, and plot polypeptide concentration on the X-axis (log scale) and fluorescence or absorbance of the Y-axis (linear scale). Interpolate the concentration of the polypeptide in the sample using the standard curve.

Example 23: Formulating a Polypeptide

The secreted polypeptide composition will be formulated and dosed in a fashion consistent with good medical practice, taking into account the clinical condition of the individual patient (especially the side effects of treatment with the secreted polypeptide alone), the site of delivery, the method of administration, the scheduling of administration, and other factors known to practitioners. The "effective amount" for purposes herein is thus determined by such considerations.

As a general proposition, the total pharmaceutically effective amount of secreted polypeptide administered parenterally per dose will be in the range of about 1 µg/kg/day to 10 mg/kg/day of patient body weight, although, as noted above, this will be subject to therapeutic discretion. More preferably, this dose is at least 0.01 mg/kg/day, and most preferably for humans between about 0.01 and 1 mg/kg/day for the hormone. If given continuously, the secreted polypeptide is typically administered at a dose rate of about 1 µg/kg/hour to about 50 µg/kg/hour, either by 1-4 injections per day or by continuous subcutaneous infusions, for example, using a mini-pump. An intravenous bag solution may also be employed. The length of treatment needed to observe changes and the interval following treatment for responses to occur appears to vary depending on the desired effect.

Pharmaceutical compositions containing the secreted protein of the invention are administered orally, rectally, parenterally, intracistemally, intravaginally,

intraperitoneally, topically (as by powders, ointments, gels, drops or transdermal patch), bucally, or as an oral or nasal spray. "Pharmaceutically acceptable carrier" refers to a non-toxic solid, semisolid or liquid filler, diluent, encapsulating material or formulation auxiliary of any type. The term "parenteral" as used herein refers to modes of administration which include intravenous, intramuscular, intraperitoneal, intrasternal, subcutaneous and intraarticular injection and infusion.

The secreted polypeptide is also suitably administered by sustained-release systems. Suitable examples of sustained-release compositions include semi-permeable polymer matrices in the form of shaped articles, e.g., films, or mirocapsules. Sustained-release matrices include polylactides (U.S. Pat. No. 3,773,919, EP 58,481), 10 copolymers of L-glutamic acid and gamma-ethyl-L-glutamate (Sidman, U. et al., Biopolymers 22:547-556 (1983)), poly (2- hydroxyethyl methacrylate) (R. Langer et al., J. Biomed. Mater. Res. 15:167-277 (1981), and R. Langer, Chem. Tech. 12:98-105 (1982)), ethylene vinyl acetate (R. Langer et al.) or poly-D- (-)-3-hydroxybutyric acid (EP 133,988). Sustained-release compositions also include liposomally entrapped 15 polypeptides. Liposomes containing the secreted polypeptide are prepared by methods known per se: DE 3,218,121; Epstein et al., Proc. Natl. Acad. Sci. USA 82:3688-3692 (1985); Hwang et al., Proc. Natl. Acad. Sci. USA 77:4030-4034 (1980); EP 52,322; EP 36,676; EP 88,046; EP 143,949; EP 142,641; Japanese Pat. Appl. 83-118008; U.S. Pat. Nos. 4,485,045 and 4,544,545; and EP 102,324. Ordinarily, the liposomes 20 are of the small (about 200-800 Angstroms) unilamellar type in which the lipid content is greater than about 30 mol. percent cholesterol, the selected proportion being adjusted for the optimal secreted polypeptide therapy.

For parenteral administration, in one embodiment, the secreted polypeptide is formulated generally by mixing it at the desired degree of purity, in a unit dosage injectable form (solution, suspension, or emulsion), with a pharmaceutically acceptable carrier, i.e., one that is non-toxic to recipients at the dosages and concentrations employed and is compatible with other ingredients of the formulation. For example, the formulation preferably does not include oxidizing agents and other compounds that are known to be deleterious to polypeptides.

Generally, the formulations are prepared by contacting the polypeptide uniformly and intimately with liquid carriers or finely divided solid carriers or both. Then, if necessary, the product is shaped into the desired formulation. Preferably the carrier is a parenteral carrier, more preferably a solution that is isotonic with the blood of the recipient. Examples of such carrier vehicles include water, saline, Ringer's solution, and dextrose solution. Non-aqueous vehicles such as fixed oils and ethyl oleate are also useful herein, as well as liposomes.

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The carrier suitably contains minor amounts of additives such as substances that enhance isotonicity and chemical stability. Such materials are non-toxic to recipients at the dosages and concentrations employed, and include buffers such as phosphate, citrate, succinate, acetic acid, and other organic acids or their salts; antioxidants such as ascorbic acid; low molecular weight (less than about ten residues) polypeptides, e.g., polyarginine or tripeptides; proteins, such as serum albumin, gelatin, or immunoglobulins; hydrophilic polymers such as polyvinylpyrrolidone; amino acids, such as glycine, glutamic acid, aspartic acid, or arginine; monosaccharides, disaccharides, and other carbohydrates including cellulose or its derivatives, glucose, manose, or dextrins; chelating agents such as EDTA; sugar alcohols such as mannitol or sorbitol; counterions such as sodium; and/or nonionic surfactants such as polysorbates, poloxamers, or PEG.

The secreted polypeptide is typically formulated in such vehicles at a concentration of about 0.1 mg/ml to 100 mg/ml, preferably 1-10 mg/ml, at a pH of about 3 to 8. It will be understood that the use of certain of the foregoing excipients, carriers, or stabilizers will result in the formation of polypeptide salts.

Any polypeptide to be used for therapeutic administration can be sterile. Sterility is readily accomplished by filtration through sterile filtration membranes (e.g., 0.2 micron membranes). Therapeutic polypeptide compositions generally are placed into a container having a sterile access port, for example, an intravenous solution bag or vial having a stopper pierceable by a hypodermic injection needle.

Polypeptides ordinarily will be stored in unit or multi-dose containers, for example, sealed ampoules or vials, as an aqueous solution or as a lyophilized formulation for reconstitution. As an example of a lyophilized formulation, 10-ml vials are filled with 5 ml of sterile-filtered 1% (w/v) aqueous polypeptide solution, and the resulting mixture is lyophilized. The infusion solution is prepared by reconstituting the lyophilized polypeptide using bacteriostatic Water-for-Injection.

The invention also provides a pharmaceutical pack or kit comprising one or more containers filled with one or more of the ingredients of the pharmaceutical compositions of the invention. Associated with such container(s) can be a notice in the form prescribed by a governmental agency regulating the manufacture, use or sale of pharmaceuticals or biological products, which notice reflects approval by the agency of manufacture, use or sale for human administration. In addition, the polypeptides of the present invention may be employed in conjunction with other therapeutic compounds.

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Example 24: Method of Treating Decreased Levels of the Polypeptide

It will be appreciated that conditions caused by a decrease in the standard or normal expression level of a secreted protein in an individual can be treated by administering the polypeptide of the present invention, preferably in the secreted form. Thus, the invention also provides a method of treatment of an individual in need of an increased level of the polypeptide comprising administering to such an individual a pharmaceutical composition comprising an amount of the polypeptide to increase the activity level of the polypeptide in such an individual.

For example, a patient with decreased levels of a polypeptide receives a daily dose 0.1-100 ug/kg of the polypeptide for six consecutive days. Preferably, the polypeptide is in the secreted form. The exact details of the dosing scheme, based on administration and formulation, are provided in Example 23.

Example 25: Method of Treating Increased Levels of the Polypeptide

Antisense technology is used to inhibit production of a polypeptide of the present invention. This technology is one example of a method of decreasing levels of a polypeptide, preferably a secreted form, due to a variety of etiologies, such as cancer.

For example, a patient diagnosed with abnormally increased levels of a polypeptide is administered intravenously antisense polynucleotides at 0.5, 1.0, 1.5, 2.0 and 3.0 mg/kg day for 21 days. This treatment is repeated after a 7-day rest period if the treatment was well tolerated. The formulation of the antisense polynucleotide is provided in Example 23.

Example 26: Method of Treatment Using Gene Therapy

One method of gene therapy transplants fibroblasts, which are capable of expressing a polypeptide, onto a patient. Generally, fibroblasts are obtained from a subject by skin biopsy. The resulting tissue is placed in tissue-culture medium and separated into small pieces. Small chunks of the tissue are placed on a wet surface of a tissue culture flask, approximately ten pieces are placed in each flask. The flask is turned upside down, closed tight and left at room temperature over night. After 24 hours at room temperature, the flask is inverted and the chunks of tissue remain fixed to the bottom of the flask and fresh media (e.g., Ham's F12 media, with 10% FBS, penicillin and streptomycin) is added. The flasks are then incubated at 37°C for approximately one week.

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At this time, fresh media is added and subsequently changed every several days. After an additional two weeks in culture, a monolayer of fibroblasts emerge. The monolayer is trypsinized and scaled into larger flasks.

pMV-7 (Kirschmeier, P.T. et al., DNA, 7:219-25 (1988)), flanked by the long terminal repeats of the Moloney murine sarcoma virus, is digested with EcoRI and HindIII and subsequently treated with calf intestinal phosphatase. The linear vector is fractionated on agarose gel and purified, using glass beads.

The cDNA encoding a polypeptide of the present invention can be amplified using PCR primers which correspond to the 5' and 3' end sequences respectively as set forth in Example 1. Preferably, the 5' primer contains an EcoRI site and the 3' primer includes a HindIII site. Equal quantities of the Moloney murine sarcoma virus linear backbone and the amplified EcoRI and HindIII fragment are added together, in the presence of T4 DNA ligase. The resulting mixture is maintained under conditions appropriate for ligation of the two fragments. The ligation mixture is then used to transform bacteria HB101, which are then plated onto agar containing kanamycin for the purpose of confirming that the vector has the gene of interest properly inserted.

The amphotropic pA317 or GP+am12 packaging cells are grown in tissue culture to confluent density in Dulbecco's Modified Eagles Medium (DMEM) with 10% calf serum (CS), penicillin and streptomycin. The MSV vector containing the gene is then added to the media and the packaging cells transduced with the vector. The packaging cells now produce infectious viral particles containing the gene (the packaging cells are now referred to as producer cells).

Fresh media is added to the transduced producer cells, and subsequently, the media is harvested from a 10 cm plate of confluent producer cells. The spent media, containing the infectious viral particles, is filtered through a millipore filter to remove detached producer cells and this media is then used to infect fibroblast cells. Media is removed from a sub-confluent plate of fibroblasts and quickly replaced with the media from the producer cells. This media is removed and replaced with fresh media. If the titer of virus is high, then virtually all fibroblasts will be infected and no selection is required. If the titer is very low, then it is necessary to use a retroviral vector that has a selectable marker, such as neo or his. Once the fibroblasts have been efficiently infected, the fibroblasts are analyzed to determine whether protein is produced.

The engineered fibroblasts are then transplanted onto the host, either alone or after having been grown to confluence on cytodex 3 microcarrier beads.

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It will be clear that the invention may be practiced otherwise than as particularly described in the foregoing description and examples. Numerous modifications and variations of the present invention are possible in light of the above teachings and, therefore, are within the scope of the appended claims.

The entire disclosure of each document cited (including patents, patent applications, journal articles, abstracts, laboratory manuals, books, or other disclosures) in the Background of the Invention, Detailed Description, and Examples is hereby incorporated herein by reference.

```
(1) GENERAL INFORMATION:
           (i) APPLICANT:
                            Human Genome Sciences, Inc. et al.
           (ii) TITLE OF INVENTION: 87 Human Secreted Proteins
           (iii) NUMBER OF SEQUENCES: 323
5
           (iv) CORRESPONDENCE ADDRESS:
                 (A) ADDRESSEE: Human Genome Sciences, Inc.
                 (B) STREET: 9410 Key West Avenue
                 (C) CITY: Rockville
10
                 (D) STATE: Maryland
                 (E) COUNTRY: USA
                  (F) ZIP: 20850
15
           (v) COMPUTER READABLE FORM:
                  (A) MEDIUM TYPE: Diskette, 3.50 inch, 1.4Mb storage
                  (B) COMPUTER: HP Vectra 486/33
                  (C) OPERATING SYSTEM: MSDOS version 6.2
20
                  (D) SOFTWARE: ASCII Text
            (vi) CURRENT APPLICATION DATA:
                  (A) APPLICATION NUMBER:
25
                  (B) FILING DATE: March 19, 1998
                  (C) CLASSIFICATION:
            (vii) PRIOR APPLICATION DATA:
30
                  (A) APPLICATION NUMBER:
                  (B) FILING DATE:
            (viii) ATTORNEY/AGENT INFORMATION:
35
                  (A) NAME: A. Anders Brookes
                  (B) REGISTRATION NUMBER: 36,373
                  (C) REFERENCE/DOCKET NUMBER: PZ004PCT
 40
            (vi) TELECOMMUNICATION INFORMATION:
                   (A) TELEPHONE: (301) 309-8504
                   (B) TELEFAX: (301) 309-8439
 45
      (2) INFORMATION FOR SEQ ID NO: 1:
             (i) SEQUENCE CHARACTERISTICS:
                   (A) LENGTH: 733 base pairs
 50
                   (B) TYPE: nucleic acid
                   (C) STRANDEDNESS: double
                   (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 1:
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GGGATCCGGA GCCCAAATCT TCTGACAAAA CTCACACATG CCCACCGTGC CCAGCACCTG	60
AATTCGAGGG TGCACCGTCA GTCTTCCTCT TCCCCCCAAA ACCCAAGGAC ACCCTCATGA	120
TCTCCCGGAC TCCTGAGGTC ACATGCGTGG TGGTGGACGT AAGCCACGAA GACCCTGAGG	180
TCAAGTTCAA CTGGTACGTG GACGGCGTGG AGGTGCATAA TGCCAAGACA AAGCCGCGGG	240
AGGAGCAGTA CAACAGCACG TACCGTGTGG TCAGCGTCCT CACCGTCCTG CACCAGGACT	300
GGCTGAATGG CAAGGAGTAC AAGTGCAAGG TCTCCAACAA AGCCCTCCCA ACCCCCATCG	360
AGAAAACCAT CTCCAAAGCC AAAGGGCAGC CCCGAGAACC ACAGGTGTAC ACCCTGCCCC	420
CATCCCGGGA TGAGCTGACC AAGAACCAGG TCAGCCTGAC CTGCCTGGTC AAAGGCTTCT	480
ATCCAAGCGA CATCGCCGTG GAGTGGGAGA GCAATGGGCA GCCGGAGAAC AACTACAAGA	540
CCACGCCTCC CGTGCTGGAC TCCGACGGCT CCTTCTTCCT CTACAGCAAG CTCACCGTGG	600
ACAAGAGCAG GTGGCAGCAG GGGAACGTCT TCTCATGCTC CGTGATGCAT GAGGCTCTGC	660
ACAACCACTA CACGCAGAAG AGCCTCTCCC TGTCTCCGGG TAAATGAGTG CGACGGCCGC	720
GACTCTAGAG GAT	733
	•
(2) INFORMATION FOR SEO ID NO: 2:	
-	
(A) LENGTH: 5 amino acids	
(D) TOPOLOGY: linear	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 2:	
Trp Ser Xaa Trp Ser	
(2) INFORMATION FOR SEQ ID NO: 3:	
(i) SEQUENCE CHARACTERISTICS:	
(B) TYPE: nucleic acid	
(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 3:	
COCCOMOCA A MINEROCCOA A A MOCA CAMPA MARCALA A A MARCALA A MARCALA A A MARCALA A A MARCALA A A MARCALA A MA	
GCGCCTCGAG ATTTCCCCGA AATCTAGATT TCCCCGAAAT GATTTCCCCG AAATGATTTC	60
	AATTCCAGGG TGCACCGTCA GTCTTCCTCT TCCCCCCAAA ACCCAAGGAC ACCCTCATGA TCTCCCGGAC TCCTGAGGTC ACATGCGTGG TGGTGGACGT AAGCCACGAA GACCCTGAGG TCAAGTTCAA CTGGTACGTG GACGGCGTGG AGGTGCATAA TGCCAAGACA AAGCCGCGGG AGGAGCAGTA CAACAGCACG TACCGTGTGG TCAGCGTCCT CACCGTCCTG CACCAGGACT GGCTGAATGG CAAGGAGTAC AAGTGCAAGG TCTCCAACAA AGCCCTCCCA ACCCCCATCG AGAAAAACCAT CTCCAAAGCC AAAGGGCAGC CCCGAGAACC ACAGGTGTAC ACCCTGCCCC CATCCCGGGA TGAGCTGACC AAAGACCAGG TCAGCCTGAC CTGCCTGGTC AAAGGCTTCT ATCCAAGGCA CATCGCCGTG GAGTGGGAGA GCAATGGGCA GCCGGAGAAC AACTACAAGA CCACGCCTCC CGTGCTGGAC TCCGACGGCT CCTTCTTCCT CTACAGCAAG CTCACCGTGG ACAAGAGCAG GTGGCAGCAG GGGAACGTCT TCTCATGCTC CGTGATGCAT GAGGCTTCC ACAACCACTA CACGCAGAAG AGCCTCTCCC TGTCTCCGGG TAAATGAGTG CGACGGCCGC GACTCTAGAG GAT (2) INFORMATION FOR SEQ ID NO: 2: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 5 amino acids (B) TYPE: amino acids (C) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 2: TTP SER Xaa Trp Ser 1

BNSDOCID: <WO 9842738A1>

	(2) INFORMATION FOR SEQ ID NO: 4:	
5	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 27 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 4:	
	GCGCCAAGCT TTTTGCAAAG CCTAGGC	27
15		
	(2) INFORMATION FOR SEQ ID NO: 5:	
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 271 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 5:	
	CTCGAGATTT CCCCGAAATC TAGATTTCCC CGAAATGATT TCCCCGAAAT GATTTCCCCG	60
20	AAATATCTGC CATCTCAATT AGTCAGCAAC CATAGTCCCG CCCCTAACTC CGCCCATCCC	120
30	GCCCCTAACT CCGCCCAGTT CCGCCCATTC TCCGCCCCAT GGCTGACTAA TTTTTTTTAT	180
	TTATGCAGAG GCCGAGGCCG CCTCGGCCTC TGAGCTATTC CAGAAGTAGT GAGGAGGCTT	240
35	TTTTGGAGGC CTAGGCTTTT GCAAAAAGCT T	271
40	(2) INFORMATION FOR SEQ ID NO: 6:	
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 32 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 6:	
50	GCGCTCGAGG GATGACAGCG ATAGAACCCC GG	32
55	(2) INFORMATION FOR SEQ ID NO: 7:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 31 base pairs	
60	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	

	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 7:	
5	GCGAAGCTTC GCGACTCCCC GGATCCGCCT C	3:
10	(2) INFORMATION FOR SEQ ID NO: 8:	
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 12 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 8:	
20	GGGGACTTTC CC	12
25	(2) INFORMATION FOR SEQ ID NO: 9:	
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 73 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 9:	
	GCGGCCTCGA GGGGACTTTC CCGGGGACTT TCCGGGGACT TTCCATCCTG	60
40	CCATCTCAAT TAG	7 3
	(2) INFORMATION FOR SEQ ID NO: 10:	
45	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 256 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double	
50	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 10:	
	CTCGAGGGGA CTTTCCCGGG GACTTTCCGG GGACTTTCCA TCTGCCATCT	60
55	CAATTAGTCA GCAACCATAG TCCCGCCCCT AACTCCGCCC ATCCCGCCCC TAACTCCGCC	120
	CAGTTCCGCC CATTCTCCGC CCCATGGCTG ACTAATTTTT TTTATTTATG CAGAGGCCGA	180
60	GGCCGCCTCG GCCTCTGAGC TATTCCAGAA GTAGTGAGGA GGCTTTTTTG GAGGCCTAGG	240

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CTTTTGCAAA AAGCTT	256
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(2) INFORMATION FOR SEQ ID NO: 11:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1679 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 11:

15 GCAGCGCACC CGGGCGATCG CTTCACGGAT GCGGACGACG TAGCCATCCT TACCTACGTG 60 AAGGAAAATG CCCGCTCGCC CAGCTCCGTC ACCGGTAACG CCTTGTGGAA AGCGATGGAG 120 AAGAGCTCGC TCACGCAGCA CTCGTGGCAG TCCCTGAAGG ACCGCTACCT CAAGCACCTG 180 20 CGGGGCCAGG AGCATAAGTA CCTGCTGGGG GACGCGCCGG TGAGCCCCTC CTCCCAGAAG 240 CTCAAGCGGA AGGCGGAGGA GGACCCGGAG GCCGCGGATA GCGGGGAACC ACAGAATAAG 300 25 AGAACTCCAG ATTTGCCTGA AGAAGAGTAT GTGAAGGAAG AAATCCAGGA GAATGAAGAA 360 GCAGTCAAAA AGATGCTTGT GGAAGCCACC CGGGAGTTTG AGGAGGTTGT GGTGGATGAG 420 AGCCCTCCTG ATTTTGAAAT ACATATAACT ATGTGTGATG ATGATCCACC CACACCTGAG 480 30 GAAGACTCAG AAACACAGCC TGATGAGGAG GAAGAAGAAG AAGAAGAAAA AGTTTCTCAA 540 CCAGAGGTGG GAGCTGCCAT TAAGATCATT CGGCAGTTAA TGGAGAAGTT TAACTTGGAT 600 35 CTATCAACAG TTACACAGGC CTTCCTAAAA AATAGTGGTG AGCTGGAGGC TACTTCCGCC 660 720 TTCTTAGCGT CTGGTCAGAG AGCTGATGGA TATCCCATTT GGTCCCGACA AGATGACATA GATTTGCAAA AAGATGATGA GGATACCAGA GAGGCATTGG TCAAAAAATT TGGTGCTCAG 780 40 840 AAAGTCATGG TAGGTGAGGT GGTTAAAAAA AATTGTGACC AATGAACTTT AGAGAGTTCT 900 45 TGCATTGGAA CTGGCACTTA TTTTCTGACC ATCGCTGCTG TTGCTCTGTG AGTCCTAGAT 960 TTTTGTAGCC AAGCAGAGTT GTAGAGGGGG ATAAAAAGAA AAGAAATTGG ATGTATTTAC 1020 1080 AGCTGTCCTT GAACAAGTAT CAATGTGTTT ATGAAAGGAA GATCTAAATC AGACAGGAGT 50 TGGTCTACAT AGTAGTAATC CATTGTTGGA ATGGAACCCT TGCTATAGTA GTGACAAAGT 1140 GAAAGGAAAT TTAGGAGGCA TAGGCCATTT CAGGCAGCAT AAGTAATCTC CTGTCCTTTG 1200 55 GCAGAAGCTC CTTTAGATTG GGATAGATTC CAAATAAAGA ATCTAGAAAT AGGAGAAGAT 1260 TTAATTATGA GGCCTTGAAC ACGGATTATC CCCAAACCCT TGTCATTTCC CCCAGTGAGC 1320 TCTGATTTCT AGACTGCTTT GAAAATGCTG TATTCATTTT GCTAACTTAG TATTTGGGTA 1380 60

	CCCTGCTCTT	TGGCTGTTCT	TTTTTTGGAG	CCCTTCTCAG	TCAAGTCTGC	CGGATGTCTT	1440
5	TCTTTACCTA	CCCCTCAGTT	ТТССТТАААА	CGCGCACACA	ACTCTAGAGA	GTGTTAAGAA	1500
J	TAATGTTACT	TGGTTAATGT	GTTATTTATT	GAGTATTGTT	TGTGCTAAGC	ATTGTGTTAG	1560
	ATTTAAAAAA	TTAGTGGATT	GACTCCACTT	TGTTGTGTTG	TTTTCATTGT	TGAAAATAAA	1620
10	TATAACTTTG	TATTCGAAAA	ааааааааа	AAAATNRCTG	CGGNCCGACA	AGGGAATTC	1679

15 (2) INFORMATION FOR SEQ ID NO: 12:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1830 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 12:

25	GCGACCGCGC	CCTTCAGCTA	GCTCGCTCGC	TCGCTCTGCT	TCCCTGCTGC	CGGCTGCGCA	60
	TGGCTTNGGC	GTTGGCGGCG	CTGGCGGCGG	TCGAGCNGCC	TGCGSAGCCG	GTACCAGCAG	120
30	TTGCAGAATG	AAGAAGAGTC	TGGAGAACCT	GAACAGGCTG	CAGGTGATGC	TCCTCCACCT	180
	TACAGCAGCA	TTTCTGCAGA	GAGCGCACAT	NATTTTGACT	ACAAGGATGA	GTCTGGGTTT	240
	CCAAAGCCCC	CATCTTACAA	TGTAGCTACA	ACACTGCCCA	GTTATGATGA	AGCGGAGAGG	300
35	ACCAAGGCTG	AAGCTACTAT	CCCTTTGGTT	CCTGGGAGAG	ATGAGGATTT	TGTGGGTCGG	360
	GATGATTTTG	ATGATGCTGA	CCAGCTGAGG	ATAGGAAATG	ATGGGATTTT	CATGTTAACT	420
40	TTTTTCATGG	CATTCCTCTT	TAACTGGATT	GGGTTTTTCC	TGTCTTTTTG	CCTGACCACT	480
	TCAGCTGCAG	GAAGGTATGG	GGCCATTTCA	GGATTTGGTC	TCTCTCTAAT	TAAATGGATC	540
	CTGATTGTCA	GGTTTTCCAC	CTATTTCCCT	GGATATTTTG	ATGGTCAGTA	CTGGCTCTGG	600
45	TGGGTGTTCC	TTGTTTTAGG	CTTTCTCCTG	TTTCTCAGAG	GATTTATCAA	TTATGCAAAA	660
	GTTCGGAAGA	TGCCAGAAAC	TTTCTCAAAT	CTCCCCAGGA	CCAGAGTTCT	CTTTATTTAT	720
50	TAAAGATGTT	TTCTGGCAAA	GGCCTTCCTG	CATTTATGAA	TTCTCTCTCA	AGAAGCAAGA	780
	GAACACCTGC	AGGAAGTGAA	TCAAGATGCA	GAACACAGAG	GAATAATCAC	CTGCTTTAAA	840
	AAAATAAAGT	ACTGTTGAAA	AGATCATTTC	TCTCTATTTG	TTCCTAGGTG	TAAAATTTTA	900
55	ATAGTTAATG	CAGAATTCTG	TAATCATTGA	ATCATTAGTG	GTTAATGTTT	GAAAAAGCTC	960
	TTGCAATCAA	GTCTGTGATG	TATTAATAAT	GCCTTATATA	TTGTTTGTAG	TCATTTTAAG	1020
60	TAGCATGAGC	CATGTCCCTG	TAGTCGGTAG	GGGGCAGTCT	TGCTTTATTC	ATCCTCCATC	1080

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		CTTGGAATTA	አልጥልጥጥርጥል A	САТАТСТАТА	ATGCTGGCCA	TTTTAAAGGG	1140
	TCAAAATGAA	CIIGGAAIIA	MINITOIL.	<u></u>			
	GTTTTCTCAA	AAGTTAAACT	TTTGTTATGA	CTGTGTTTTT	GCACATAATC	CATATTTGCT	1200
5	GTTCAAGTTA	ATCTAGAAAT	TTATTCAATT	CTGTATGAAC	ACCTGGAAGC	AAAATCATAG	1260
	TGCAAAAATA	CATTTAAGGT	GTGGTCAAAA	ATAAGTCTTT	AATTGGTAAA	TAATAAGCAT	1320
	TAATTTTTTA	TAGCCTGTAT	TCACAATTCT	GCGGTACCTT	ATTGTACCTA	AGGGATTCTA	1380
10	AAGGTGTTGT	CACTGTATAA	AACAGAAAGC	ACTAGGATAC	AAATGAAGCT	TAATTACTAA	1440
	AATGTAATTC	TTGACACTCT	TTCTATAATT	AGCGTTCTTC	ACCCCCACCC	CCACCCCCAC	1500
15	CCCCCTTATT	TTCCTTTTGT	CTCCTGGTGA	TTAGGCCAAA	GTCTGGGAGT	AAGGAGAGGA	1560
	TTAGGTACTT	AGGAGCAAAG	AAAGAAGTAG	CTTGGAACTT	TTGAGATGAT	CCCTAACATA	1620
20	CTGTACTACT	TGCTTTTACA	ATGTGTTAGC	AGAAACCAGT	GGGTTATAAT	GTAGAATGAT	1680
20	GTGCTTTCTG	CCCAAGTGGT	AATTCATCTT	GGTTTGCTAT	GTTAAAACTG	TAAATACAAC	1740
	AGAACATTAA	TAAATATCTC	TTGTGTAGCA	CCTTTTAAAA	AAAAAAAA	AAAAAAAA	1800
25	ААААААААА	AANCCCGGGG	GGGGGCCCCI	1			1830

30 (2) INFORMATION FOR SEQ ID NO: 13:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1212 base pairs

(B) TYPE: nucleic acid(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 13:

TGTTTGAAGT TGTTACTTTT GTTTACAGCA AAGTTTGATG TAGTGTGCAG TAGTGAGCTC 60 40 TAGACTGATC TTTTTCTAAA TCAGAAAGTG ATTAAAGTAT GCACAACCAA AGGCAGGTTT 120 TTCTTTTCA TTTATTCAGC AACTATTTAT TAAGCATCAA CTCTGTGCCA GGCACGTTAC 180 45 TAGCTGCTAC ATACTGTCTG AACATGACAT ACGGTTAAGT AACTTTACAA TTATTATCAA 240 ATACTTCAAT GTAGATATTT CTTAAGTTGA AATAGCATTA ACTAGGATAA TGCTTTCATG 300 TTATTTATT TGTCTTGTGA TAGAAATTCA ACTTTGTACC ATCTTAAAAC TAGGTTGCTA 360 50 TAAAAATAGG AGGATGAAGT CAATAAAGTT TATGCCAGTT TAAAAACTGG AAGGAAAAGG 420 TAAGAGCTCT CCATTATAAA ATAGTTGCAT TCGGTTAATT TTTACACATT AGTGCATTGC 480 55 540 GTATATCAAC TGGCCCTCAA TGAAGCATTT AAGTGCTTGG AATTTTACTA AACTGACTTT TTTGCAACTT TGGGAGATTT TTGAGGGGAG TGTTGAAAAT TGCCAAACAC TCACCTCTTA 600 CTCAAAACTT CAAATAAAAT ACACATTTTC AAGAGGGAGC ACCTTTTATA TTTGATAAGT 660 60

720

780

	TTTCATTATA AACCTTATAA TACCAGTCAC AAAGAGGTTG TCTGTCTATG GTTTAGCAAA	720
5	CATTIGCTIT TCTTTTIGGA AGTGTGATTG CAATTGCAGA ACAGAAAGTG AGAAAACACT	780
J	GCCAGCGGTG ATTGCTACTT GAGGTAGTTT TTTACAACTA CCATTTCCCC TCCATGAAAT	840
	TATGTGAAAT TTATTTTATC TTTGGGAAAA GTTGAGAAGA TAGTAAAAGA ATTAGGAATT	900
10	TAAAATTACA GGGAAAAATA TGTAAGTGAA AAGCAATAAA TATTTTGTTC ACTTTGCTAT	960
	CAAGATGTTC ACTATCAGAT ATTTATTATA TGGCAGCAAT TTATATTTTT AATCATTGCC	1020
15	CATTAATAGA CGCAGTAAAA TATTTTTGAA TCAGACATTT GGGGTTTGTA TGTGCATTAA	1080
13	AATTGTCTTT TGTACTGTAA GTTACTGTTA ATTTGAATAT TTTATTGAAC TGTCTCCCTG	1140
	TGCCTTTATA ATATAAAGTT GTTTCTACAA CTTTTAATGA TCTTAATAAA GAATACTTTA	1200
20	AGAAAAAAA AA	1212
25	(2) INFORMATION FOR SEQ ID NO: 14:	
	(i) SEQUENCE CHARACTERISTICS:	
30	(A) LENGTH: 2061 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 14:	
35	GGTTTTCCTC CGACTTCCGG ACATCTCCCT GGGAGTCGCG CAGAGTGGAG TCAAAAGGCAA	60
	CCAGTGCTCG CTGCGGTCTC TGGGGATCGG GACCGCGGCG GCGGCCCGCG AGCGGGATGT	120
40	TCCGGGGCTT GAGCAGTTGG TTGGGCTTGC AGCAGCCGGT GGCAGCCGGT GGGCAGCCCA	180
70	ATGGAGATGC TCCACCCGAG CAGCCGTCCG AGACGGTGGC TGAGTCTGCG GAGGAGGAGC	240
	TGCAGCAAGC GGGAGACCAG GAGCTCCTCC ACCAGGCCAA AGACTTCGGC AACTATTTAT	300
45	TTAACTPTGC ATCTGCTGCC ACAAAAAAGA TAACTGAATC AGTTGCTGAA ACAGCACAAA	360
	CAATAAAGAA ATCCGTAGAA GAAGGAAAAA TAGATGGCAT CATTGACAAG ACAATTATAG	420
50	GAGATTTTCA GAAGGAACAG AAAAAATTTG TTGAAGAGCA ACATACAAAG AAGTCAGAAG	480
20	CAGCTGTGCC CCCATGGGTT GACACTAACG ATGAAGAAAC AATTCAACAA CAAATTTTGG	540
	CCTTATCAGC TGACAAGAGG AATTTCCTTC GTGACCCTCC GGCTGGCGTG CAATTTAATT	600
55	TCGACTTTGA TCAGATGTAC CCCGTGGCCC TGGTCATGCT CCAGGAGGAT GAGCTGCTAR	660

CAAGATGAGA TTTGCCCTCG TTCCTAAACT TGTGAAGGAA GAAGTGTTCT GGAGGAACTA

CTTTTACCGC GTCTCCCTGA TTAAGCAGTC AGCCCAGCTC ACGCCCTGG CTGCCCAACA

120

	GCAGCCCGCA GGGAAGGGAG GAGAAGAGCA ATGGCAGAGA GCAAGATTTG CCGCTGGAGA	840
	GGCAGTACGG CCCAAAACGC CACCCGTTGT AATCAAATCT CAGCTTAAAA CTCAAGAGGA	900
5	TGAGGAAGAA ATTTCTACTA GCCCAGGTGT TTCTGAGTTT GTCAGTGATG CCTTCGATGC	960
	CTGTAACCTA AATCAGGAAG ATCTAAGGAA AGAAATGGAG CAACTAGTGC TTGACAAAAA	1020
10	GCAAGAGGAG ACAGCCGTAC TGGAAGAGGA TTCTGCAGAT TGGGAAAAAG AACTGCAGCA	1080
10	GGAACTTCAA GAATATGAAG TGGTGACAGA ATCTGAAAAA CGAGATGAAA ACTGGGATAA	1140
	GGAAATAGAG AAAATGCTTC AAGAGGAAAA TTAGCTGTTC CTGAAATAGA AGAATAATCC	1200
15	TTAACAGTCT GCAAACTGAC ATTAAATTCT AGATGTTGAC AATTACTGAA TCAGAAGGCA	1260
	TGAAAGAGTA TAATTTTATG AAATTCAAAA TTATTCTTTT TTCAAGTTGA AACTTGCCTC	1320
20	TTCTACTTTA AAAAAGTATA TAGAACAGTT ACTTCTAATA ATCAGAAAGA GATGTTTTAT	1380
20	AGAACATTTC TTTAATATAA AGTTAGAGAT GTCTTCATAG GCAGTATGGC TATCTTTGCC	1440
	ACAGAAACAT AAGTAAAATT TTAGAGTTCT GTTTTCCATG AGGTCAAAAA TATAATTTAT	1500
25	TCCTCAGTCA TGGTTTTCTA AATATCTGTA CTCCACATTC CATTTTAATT GATATGAGGG	1560
	TGTTAAAGTA CCTACTTAAT GGGTTGATTA CTATCAAAAT GACCAAATTA TACCAAAGAA	1620
30	CTTAAGAGGA AGCACTTTCA GAACTATTCA CTTGCCAGGT ATTTTCTAAA ATTCCACCTG	1680
30	AAAGCCAAAA GATAAAATAC ATNAGTTGGA TITTAATGAT ATAAGCATCA CACAATTTTA	1740
	CATTAAGAAA TACTGTGCAG CCCATGCGTG GTGGCTCAGG CCTGTAATCC CAGCANTTTG	1800
35	GGAGGCCGAG GTGGGCAGAT CACCGGAGGT CAGGAGTTCG AGACCAGCCT TGCCAACATA	1860
	GTGAAACCCT GTCTTTACTA AAAATACAAA AATTAGCCGG GCATGGTGGC AGGCACCTGT	1920
40	AATCCCAGCT ACTAGGGAGG CTTTTGAACC CAGGAGGCAG AGGTTGCAGC GAGCTGAGAT	1980
70	CGCGCCACTG CACTCCAGCC TGGGTGATAG AGTGAGATTC AGTCTCAAAA AAAAAAAAAA	2040
	AAAAAAAAA AATGACCTCG A	2061
45		
	(2) INFORMATION FOR SEQ ID NO: 15:	
50	(i) SEOUENCE CHARACTERISTICS:	
30	(A) LENGTH: 1412 base pairs	
	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
55	(D) TOPOLOGY: linear	
33	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 15:	

CCCTTCATCT GCGTTGCCAG GAACCCTGTC AGCAGAAACT TCTCAAGCCC CATCCTTGCC

AGGAAGCTCT GTGAAGGTGC TGCTGATGAC CCAGATTCCT CCATGGTCCT CCTGTGTCTC

	CTGTTGGTGC	CCCTCCTGCT	CAGTCTCTTT	GTACTGGGGC	TATTTCTTTG	GTTTCTGAAG	180
5	AGAGAGAGAC	AAGAAGAGTA	CATTGAAGAG	AAGAAGAGAG	TGGACATTTG	TCGGGAAACT	240
	CCTAACATAT	GCCCCCATTC	TGGAGAGAAC	ACAGAGTACG	ACACAATCCC	TCACACTAAT	300
	AGAACAATCC	TAAAGGAAGA	TCCAGCAAAT	ACGGTTTACT	CCACTGTGGA	AATACCGAAA	360
10	AAGATGGAAA	ATCCCCACTC	ACTGCTCACG	ATGCCAGACA	CACCAAGGCT	ATTTGCCTAT	420
	GAGAATGTTA	TCTAGACAGC	AGTGCACTCC	CCTAAGTCTC	TGCTCAAAAA	AAAAACAATT	480
15	CTCGGCCCAA	AGAAAACAAT	CAGAAGAATT	CACTGATTTG	ACTAGAAACA	TCAAGGAAGA	540
	ATGAAGAACG	TTGACTTTTT	TCCAGGATAA	ATTATCTCTG	ATGCTTCTTT	AGATTTAAGA	600
	GTTCATAATT	CCATCCACTG	CTGAGAAATC	TCCTCAAACC	CAGAAGGTTT	AATCACTTCA	660
20	TCCCAAAAAT	GGGATTGTGA	ATGTCAGCAA	АССАТААААА	AAGTGCTTAG	AAGTATTCCT	720
	ATAAAAATGT	AAATGCAAGG	TCACACATAT	TAATGACAGC	CTGTTGTATT	AATGATGGCT	780
25	CCAGGTCAGT	GTCTGGAGTT	TCATTCCATC	CCAGGGCTTG	GATGTCAGGA	TTATACCAAG	840
	AGTCTTGCTA	CCAGGAGGGC	AAGAAGACCA	AAACAGACAG	ACAAGTCCAG	CAGAAGCAGA	900
	TGCACCTGAC	AAAAATGGAT	GTATTAATTG	GCTCTATAAA	CTATGTGCCC	AGCAYTATGC	960
30	TGAGCTTACA	CTAATTGGTC	AGACATGCTG	TCTGCCCTCA	TGAAATTGGC	TCCAAATGAW	1020
	TGAACTACTT	TCATGAGCAG	TTGTAGCAGG	CCTGACCACA	GATTCCCAGA	GGGCCAGGTG	1080
35	TGGATCCACA	GGACTTGAAG	GTCAAAGTTC	ACAAAGATGA	AGAATCAGGG	TAGCTGACCA	1140
	TGTTTGGCAG	ATACTATAAT	GGAGACACAG	AAGTGTGCAT	GGCCCAAGGA	CAAGGACCTC	1200
	CAGCCAGGCT	TCATTTATGC	ACTTGTCTGC	AAAAGAAAAG	TCTAGGTTTT	AAGGCTGTGC	1260
40	CAGAACCCAT	CCCAATAAAG	AGACCGAGTC	TGAAGTCACA	TTGTAAATCT	AGTGTAGGAG	1320
	ACTTGGAGTC	AGGCAGTGAG	ACTGGTGGGG	CACGGGGGGC	ANTGGGTANT	GTAAACCTTT	1380
1 5	TAAAGATGGT	TAATTCNTCA	TTAGTGTTTT	TT			1412
-0	(2) INFORMA	TION FOR SE	Q ID NO: 16	i:			
50	(i)	SEQUENCE CH	IARACTERISTI	CS:			
		(B) TYPE	FTH: 1052 ba E: nucleic a	acid			
55			ANDEDNESS: 0 DLOGY: linea				

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 16:

TTCCTCTCCT CTCTCTACCC CTCCTGTCTC TCCTCCCCTC CTCTCTCTTC CTCTCCTCTC

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	TCTCTTCCTC	TCCTCTCTCT	TCCCTTCCTG	TCTCTCTTCC	CCTCCTCTCT	CTCTTCCTGT	120
	CCTCTATCTC	TTCCCCTCCT	CTATCTCTTC	СТСТССТСТС	TCTCTTCCTC	TCCTCTCTCT	180
5	CTCTTSCTTT	CTTCTCTCTC	TCCTGTCTCG	GCTGTTGTGG	GTTGCAGGTT	GGGTGCTGCT	240
	GTTGTGGTCC	TTCCCAGAAA	CTGCCAGTAG	AGGGCAGCCT	GGGCATCCTA	ATGCTTACTC	300
	TGGTTGTTAC	ACAAAGAAAA	TATTGGGGTC	ACTGGCGAGC	CCACCCACAC	TCACCAGAAT	360
10.∕	CTCCACTGTA	GTCCCCCTAA	CAAACAGCCC	TTCACTTCCT	CTCCCACTTC	AGCAATTTGT	420
	ATTTTGATGC	CATTGGCCTC	AGATCAGAGT	GTTTTAAATC	ATCACGCCCT	GGCTTATCCC	480
15	TGGTCGAGCC	AGGACACGGG	GTGCTTCAGT	GGGTCTGTCA	CCCTCTCTCC	TTGAAGCATG	540
	TIGCTTTTAT	TTATTTACTT	TTACTCTCAC	CCTGCTCCTG	TACCAGCAGG	GGCCACTTCA	600
20	AAGCCAAGGT	ACAGGGTGAT	AACTTGTGGT	CCAGCATCAG	TTTTCTCCAC	TTCTTTCTCC	660
20	CACTCACCCC	CAGCAAGGTG	CCTGGGGAGA	CTTGAGCAGA	TGTTTCATT	TGGCCTGGCC	720
	AGTGGCTGAA	AGCAGGCCTC	CAATGCACTG	TGACCTCTGG	CTTCCCCAGC	AGCTTTCCCA	780
25	GAGAGGCAGA	GGGGCCTTCC	ACAGCCCGGG	TTCTCCTGCT	GCCTCCTGCC	TGCTGCAGCT	840
	GCAGGCATTC	TGAGGGGCAA	CGTGGAGGAA	GGGCCAGGGA	TGCATGGGAT	TTTAATTGTT	900
30	TCATCACACC	TTCCCCGTGG	CAAAGAAACA	GTCAGTCCTC	TTCAGGTGTC	TTCTGGATTT	960
50	CTGGTGATGG	ACAGAGAAAT	CTTTTTACAC	TTTCAAATTA	A TGTTCAACA	ATAAAAATTG	1020
	CATTTTTAT	TTTGGAAAA	AAAAAAAA	A AA			1052
35							

(2) INFORMATION FOR SEQ ID NO: 17:

40

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 683 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

45

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 17:

AATTCGGCAG AGGCACTTAT CATGTACATA TAGCCTGTTT TTTAGCATTG TTAGACAAAG 60 TAGGCATATT CCTTTCCATC CAAGAACTCA TAACCTAGTA ATTGTAGTTG GCTGATAGCT 120 50 CATTGCCCAT ACACAAGGAT CTAACACAAC CTCTTGAATA AACATCCCCC TTATTCAGAA 180 ATGCCTTTC CTATTTCCAT ATTGCAACTT TGCTTACAAA TTTCCAATCT GTCTTTCTGT 240 55 300 TTACAGAAGA TATACAAAAT TCCTTTTGTA TGATCTCTTT ATATCTCTTG ATTTTCTTTT GTGTTTGCTA CCAAAGGGCC TGCACATAGT GAGAAGATTG TGCATGATCT GTGAGCTCTA 360 CCACACCTGG AATTAGGGAT CACCAATATG AGAAAAAAA TTGGAGGTAC AAATAACATT 420 60

	ATCATATGTW ATTGGCATAT AAATTACAGA TGTWTCTATG ACTAAAAAACC CTGTGGATAT	480
5	WAACCMAATG CAGATAAWIW TAATAAAATW TWTAAAAATW TWATCMAATA ATGATAGTGC	540
,	TATTCAAATA CTTCAAATTT GCACAGTGAT TTATTTCTTA AAATATGTTA ACACATGTGA	600
	GCCAATACAC TGAGGTCACT GGATAAATAA ACAGATTCTT GCAAAAAAAA AAAAAAAAAA	660
10	ACTCGAGGGG GGCCCGTACC CTT	683
15	(2) INFORMATION FOR SEQ ID NO: 18:	
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 1054 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 18:	
25	AAACTCATTT AGGTGACACT ATAGAAGGTA CGCCTGCAGG TACCGGTCCG GAATTCCCGG	60
	GTCGACCCAC GMGNCCGGCG ACAAGATGGC AGCAGCGTGT CGGAGCGTGA AGGGCCTGGT	120
30	GGCGGTAATA ACCGGAGGAG CCTCGGGCCT GGGCCTGGCC ACGGCGGACG ACTTGTGGGG	180
50	CAGGGAGCCT CTGCTGTGCT TCTGGACCTG CCCAACTCGG GTGGGGAGGC CCAAGCCAAG	240
	AAGTTAGGAA ACAACTGCGT TTTCGCCCCA GCCGACGTGA CCTCTGAGAA GGATGTGCAA	300
35	ACAGCTCTGG CTCTAGCAAA AGGAAAGTTT GGCCGTGTGG ATGTAGCTGT CAACTGTGCA	360
	GGCATCGCGG TGGCTAGCAA GACGTACAAC TTAAAGAAGG GCCAGACCCA TACCTTGGAA	420
40	GACTTCCAGC GAGTTCTTGA TGTGAATCTC ATGGGCACCT TCAATGTGAT CCGCCTGGTG	480
	GCTGGTGAGA TGGGCCAGAA TGAACCAGAC CAGGGAGGCC AACGTGGGGT CATCATCAAC	540
	ACTGCCAGTG TGGCTGCCTT CGAGGGTCAG GTTGGACAAG CTGCATACTC TGCTTCCAAG	600
45	GGGGGAATAG TGGGCATGAC ACTGCCCATT GCTCGGGATC TGGCTCCCAT AGGTATCCGG	660
	GTGATGACCA TTGCCCCAGG TCTGTTTGGC ACCCCACTGC TGACCAGCCT CCCAGAGAAA	720
50	GTGTGCAACT TCTTGGCCAG CCAAGTGCCC TTCCCTAGCC GACTGGGTGA CCCTGCTGAG	780
	TATGCTCACC TCGTACAGGC CATCATCGAG AACCCATTCC TCAATGGAGA GGTCATCCGG	840
	CTGGATGGGG CCATTCGTAT GCAGCCTTGA AGGGAGAAGG CAGAGAAAAC ACACGCTCCT	900
55	CTGCCCTTCC TTTCCCTGGG GTACTACTCT CCAGCTTGGG AGGAAGCCCA GTAGCCATTT	960
	TGTAACTGCC TACCAGTCGC CCTCTGTGCC TAATAAAGTC TCTTTTTCTC ACANAAAAAA	1020
60	AAAA AAAAAAAA AAAAAAAAA AAAAAAAAAA	1054

(2)	INFORMATION	FOR	SEQ	ID	NO:	19:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1393 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

10

5

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 19:

	GGAACAAGCT GGGATATGTG AGCGTTAAGC TACTCACATC CTTCAAAAAAG GTGAAACATC	60
15	TTACACGGGA CTGGAGAACC ACAGCACATG CTTTGAAGTA TTCAGTGGTC CTTGAGTTGA	120
	ATGAGGNCCA CCGGAAGGTG AGGAGGACCA CCCCCGTCCC ACTGTTCCCC AACGAGAACC	180
20	TCCCCAGCAA GATGCTCCTG GTCTATGATC TCTACTTGTY TCCTAAGCTG TGGGCTCTGG	240
	CCACCCCCA GAAGAATGGG AAGGGTGCAA GARAAGGTGA TGGAACACCT GCTCAAGCTT	300
25	TTTGGGACTT TTGGAGTCAT CTCATCAGTG CGGATCCTCA AACCTGGGAG AGAGCTGCCC	360
25	CCTGACATCC GGAGGNTCCA GCAGCCGCTA CAGCTCCTCT GACCCCGAGA GCAACCCCAC	420
	ATCCCCTATG GCGGGCCGAC GGCACGNGKC CACCAACAAG CTCAGCCCGT CTGGCCACCA	480
30	GAATCTCTTT CTGAGTCCAA ATGCCTCCCC GTGCACAAGT CCTTGGAGCA GCCCCTTGGC	540
	CCAACGCAAA GGCGTTTCCA GAAAGTCCCC ACTGGCGGAG GAAGGTAGAC TGAACTGCAG	600
25	CACCAGCCCT GAGATCTTCC GCAAGTGTAT GGATTATTCC TCTGACAGCA GCGTCACTCC	660
35	CTCTGGCAGC CCCTGGGTCC GGAGGCGTCG CCAAGCCGAG ATGGGGACCC AGGAGAAAAG	720
	CCCCGGTACG AGTCCCCTGC TCTCCCGGAA GATGCAGACT GCAGATGGGS TACCCGTAGG	780
40	TNGCTTGAGG TTGCCCAGGG GTCCTGACAA CACCAGAGGA TTTCATGGCC ATGAGAGGAG	840
	CAGGGCCTGT GTATAAATAC CTTCTATTTT TAATACAAGC TCCACTGAAA ACCACCTTCG	900
45	TTTTCAAGGT TCTGACAAAC ACCTGGCATG ACAGAATGGA ATTCGTTCCC CTTTGAGAGA	960
45	TTTTTTATTC ATGTAGACCT CTTAATTTAT CTATCTGTAA TATACATAAA TCGGTACGCC	1020
	ATGGTTTGAA GACCACCTTC TAGTTCAGGA CTCCTGTTCT TCCCAGCATG GCCACTATTT	1080
50	TGATGATGGC TGATGTGTGT GAGTGTGATG GCCCTGAAGG GCTGTAGGAC GGAGGTTCCC	1140
	TGGGGGAAGT CTGTTCTTTG GTATGGAATT TTTCTCTCTT CTTTGGTATG GAATTTTTCC	1200
	CTTCAGTGAC TGAGCTGTCC TCGATAGGCC ATGCAAGGGC TTCCTGAGAG TTCAGGAAAG	1260
55	TTCTCTTGTG CAACAGCAAG TAGCTAAGCC TATAGCATGG TGTCTTGTAG GACCAAATCG	1320
	ATGTTACCTG TCAAGTAAAT AAATAATAAA ACACCCAACT GGGAGTGCTG AAAAAAAAANA	1380
60	ANNAAAAAC TCG	1393

5	(2)	INFORMATION	FOR	SEO	TD	NO ·	20.
_	12/	TIM OLGENIA	LOK	ياد	\mathbf{L}	110:	20:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1215 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 20:

15	3CC 3 3 3 3CM	MACCINIA NAME	C111CCCCCC	1 CTC 1 CC			
13	AGGAAAAG I I	TICCNAATIG	GAAAGCGGGC	AGTGAGCGCA	ACGCAATTAA	TGTGAGTTAG	60
	NTCANTCATT	AGGCACCCCA	GGCTTTACAC	TTTATGCTTC	CGGNTCGTAT	GTTGTGTGGA	120
20	ATTGTGAGCG	GATAACAATT	TCACACAGGA	AACAGCTATG	ACCATGATTA	CGCCAAGCTN	180
	TAATACGACT	CACTATAGGG	AAAGCTGGTA	CGCCTGCAGG	TACCGGTCCG	GAATTCCCGG	240
	GTCGACCCAC	GCGTCCGCCC	ACGCGTCCGT	GAAAATCCGA	AGTGCCGCGG	AAAGTGGAGG	300
25	TGAGGGCCGC	CCGCCCTAGA	GGTGCCCGTC	CGAGAGGCAG	AGCTGACAAG	GAAGGTTTCG	360
	AGCGTTTTGC	TGGCAAAGGG	ATTTCTTACA	ACCTCCAGGC	ATGCGTCTTT	CTGCCCTGCT	420
30	GGCCTTGGCA	TCCAAGGTCA	CTCTGCCCCC	CCATTACCGC	TATGGGATGA	GCCCCCAGG	480
	CTCTGTTGCA	GACAAGAGGA	AGAACCCCCC	ATGGATCAGG	CGGCGCCCAG	TGGTTGTGGA	540
	ACCCATCTCT	GATGAAGACT	GGTATCTGTT	CTGTGGGGAC	ACGGTGGAGA	TCCTAGAAGG	600
35	CAAGGATGCC	GGGAAGCAGG	GCAAAGTGGT	TCAAGTTATC	CGGCAGCGAA	ACTGGGTGGT	660
	CGTGGGAGGG	CTGAACACAC	ATTACCGCTA	CATTGGCAAG	ACCATGGATT	ACCGGGGAAC	720
40	CATGATCCCT	AGTGAAGCCC	CCTTGCTCCA	CCGCCAGGTC	AAACTTGTGG	ATCCTATGGA	780
	CAGGAAACCC	ACTGAGATCG	AGTGGAGATT	TACTGAAGCA	GGAGAGCGGG	TACGAGTCTC	840
	CACACGATCA	GGGAGAATTA	TCCCTAAACC	CGAATTTCCC	AGAGCTGATG	GCATCGTCCC	900
45	TGAAACGTGG	ATTGATGGCC	CCAAAGACAC	ATCAGTGGAA	GATGCTTTAG	AAAGAACCTA	960
	TGTGCCCTGT	CTAAAGACAC	TGCAGGAGGA	GGTGATGGAG	GCCATGGGGA	TCAAGGAGAC	1020
50	CCGGAAATAC	AAGAAGGTCT	ATTGGTATTG	AGCCTGGGGC	AGAGCAGCTC	CTCCCCAACT	1080
50	TCTGTCCCAG	CCTTGAAGGC	TGAGGCACTT	CTTTTTCAGA	TGCCAATAAA	GAGCACTTTA	1140
	TGAGTCCTCC	АААААААА	АААААААА	АААААААА	АААААААА	АААААААА	1200
55	AAAAGGGGCG	GCCGC					1215

60 (2) INFORMATION FOR SEQ ID NO: 21:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2042 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 21:

10	CTGCATCCAG GCGCAGAATA ACCTGGGTAT CTTGTGGTCT GAAAGAGAGA AATTGAAACT	60
	GCACAGGCTT ACCTAGAGTC ATCAGAAGCA CTATATAATC AGTATATGAA AGAGGTTGGG	120
	AGTCCTCCTC TTGATCCTAC TGAGCGTTTT CTTCTGAAGA AGAGAAACTT ACTGAACAAG	180
15	AGAGATCAAA AAGATTTGAA AAGGTTTATA CTCATAACCT ATATTACCTA GCTCAAGTCT	240
	ACCAGCATCT GGAAATGTTT GAGAAGGCTG CTCACTATTG CCATAGTACA CTAAAACGCC	300
20	AGCTTGAGCA CAATGCCTAC CATCCTATAG AGTGGGCTAT CAATGCTGCT ACCTTGTCAC	360
	AGTTTTACAT CAATAAGCTA TGCTTTATGG AGGCCAGGCA CTGTTTATCA GCTGCTAATG	420
25	TCATTTTTGG TCAAACTGGA AAGATCTCAG CCACAGAAGA CACTCCTGAA GCTGAAGGAG	480
25	AAGTGCCAGA GCTTTATCAT CAAAGAAAGG GGGAAATAGC AAGGTGCTGG ATCAAATACT	540
	GTTTGACTCT CATGCAGAAT GCCCAACTCT CCATGCAGGA CAACATAGGA GAGCTTGATC	600
30	TTGATAAACA GTCTGAACTT AGAGCTTTAA GGAAAAAAGA ACTAGATGAG GAGGAAAGCA	660
	TTCGGAAAAA AGCTGTGCAG TTTGGAACCG GTGAACTGTG TGATGCCATC TCTGCAGTAG	720
25	AAGAGAAAGT GAGCTACTTG AGACCTTTAG ATTTTGAAGA AGCCAGAGAA CTTTTCTTAT	780
35	TGGGTCAGCA CTATGTCTTT GAGGCAAAAG AGTTCTTTCA GATTGATGGT TATGTCACTG	840
	ACCATATTGA AGTTGTCCAA GACCACAGTG CTCTGTTTAA GGTGCTTGCA TTCTTTGAAA	900
40	CTGACATGGA GAGACGGTGC AAGATGCATA AACGCRGAAT AGCCATGCTA GAGCCCCTAA	960
	CTGTAGACCT GAATCCACAG TATTATCTGT TGGTCAACAG ACAGATCCAG TTTGAAATTG	1020
4.5	CACATGCTTA CTATGATATG ATGGATTTGA AGGTTGCCAT TGCTGACAGG CTAAGGGATC	1080
45	CTGATTCACA CATTGTAAAA AAAATAAATA ATCTTAATAA GTCAGCACTG AAGTACTACC	1140
	AGCTCTTCTT AGACTCCCTG AGAGACCCAA ATAAAGTATT CCCTGAGCAT ATAGGGGAAG	1200
50	ATGTTCTTCG CCCTGCCATG TTAGCTAAGT TTCGAGTTGC CCGTCTCTAT GGCAAAATCA	1260
	TTACTGCAGA TCCCAAGAAA GAGCTGGAAA ATTTGGCAAC ATCATTGGGA ACATTACAAA	1320
	TTTATTGTTG ATTACTGTGA AAAGCATCCT GAGGCCGCCC AGGAAATAGA AGTTGAGCTA	1380
55	GAACTTAGTA AAGAGATGGT TAGTCTTCTC CCAACAAAAA TGGAGAGATT CAGAACCAAG	1440
	ATGGCCCTGA CTTAATCCTT GTTTTTAAAG AAAGGAAATG TGCAATATTG AAGTGATCTT	1500
60	TTTCCCTAGT CAGACAGGCC CAATTCCATT GTGATGTTTA CCTTTATAGC CAGGTGAGTG	1560

	CAGTTTGAAC TTGAGATACA GTCAACTGAG TGTTTGCTAG GATCCTAAGG AACATAAAGT	1620
5	TAATTAAAAA CTTACACCTA ATTATGTAAA TTGCCTTGTT AAAGACATGT GATTTGTATT	1680
	TTAGATGCTT GTTTCCTATT AAAATACAGA CATTTCTACC CTCAGTTTCT AAATGTAGAC	1740
	TATTTGTTGG CTAGTACTTG ATAGATTCCT TGTAAGAAAA AATGCTGGGT AATGTACCTG	1800
10	GTAACAAGCC TGTTAATATA TTAAGATTGA AAAAGTAACT TCTATAGTTA CTCCTTCTAA	1860
	AATATTIGAC TTCCTACATT CCCCCCACCC AAAATCTTTC CCTTTTGAAA ATACTAAAAA	1920
15	CTAAGTTATG TTATTATAAA GTGTAAAATG GTTTGTCTTA ATTATAGGAG AAAAAGGCCT	1980
10	TGTTAGAAAT AAAATAAACT GACTTATTTC ACTAATGAAA AAAAAAAAAA	2040
	TT	2042
20		
25	(2) INFORMATION FOR SEQ ID NO: 22: (i) SEQUENCE CHARACTERISTICS:	
30	(A) LENGTH: 1872 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 22:	
	GGGTCGACCC ACGCGTCCGA TTGGCCTAGA GCTCCTGTGA CCGAGAGCGC CACGGAAGCC	60
35	TGGGGATGAT GTCGGGCAGC TTTATTCTTT GCTTGGCTTT GGTAACTAGG TGGTCCCCTC	120
	AAGCATCCTC AGTTCCTCTT GCTGTTTATG AATCTAAGAC AAGGAAGTCC TATAGAAGCC	180
40	AAAGGGACAG GGACGGAAAG GACAGGTCCC AAGGGATGGG GCTGTCTTTA CTTGTGGAAA	240
	CCAGGAAATT GCTCCTCTCA GCCAACCAAG GTTGACCACA CACCACCCTT CCGGAGCAGC	300
	TCAGTCAGCC CTCGGGGACG RGAAACCACA AGCGCAGAGA CGCTGAGGCC CAGGCAGGTG	360
45	AAGAGGAAGT GGCTTTGGGT TTTTAAAGTA GGTGAGCGTG ACCTCTCTGA CTGCTTCTTC	420
	CCCGGGGGGG ACTGCAAACC GCTCAGGGTT GCGGCAGAGC CATGGACTTC CGGTCCCTGC	480
50	AACGGGTGAC CTAAGCGTGG TGCACCCATC AGTCACGCAG GAGGACTGAC TTGACAGACG	540
	AAAGACAAGC CCGGATGACA CAGGGTGAGA AGAGTCAGGG CCGCACCTCT GTCCCTGCAA	600
	ACCAACAGGT GCATGGTGAG TGTGGCAGTC CCCACAGCTC CACAATGGGC TCCCCCGCCA	660
55	ACGGGGACGA CAGGGATCTT CAGGAACTTC TGACCTCACC AAGTCAAGTG GACCACTCTC	720
	CACTCCACGA GGATGTGAAA CGGTTCTTTA AAATGGGATT TTAGAGCCTC GGGAATGCAT	780
60	GTGCGTCGCA TCTTTCATAT TATGGGTCAG GATAGATTCA TTTCTTGCAA CATAGTGGAA	840

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	AAGATATAAG	CTGCAGTAAT	TTGCTCTTTG	AATGACCGTC	ACCCCCAGTA	TAGGATATGC	900
	TTGTATCCCC	CCGTCACTCC	TCCGCCTGTT	TTTTAAACTT	TTCCACCACC	TGCGTCCAAA	960
5	AAGAATGTTA	TAGCGAGTGC	TCTTAAATGT	TGAACCTGGG	TGTTGCTTCC	GGGCCAGTCT	1020
	GCGTGGCTCC	ATGAAAAGCT	CACTGCTGCC	CCAGCCGGGC	TTCTTAGAGG	AGGTCAGTTG	1080
10	TCCTATGTAT	CATCATTTAC	TCTGGGAATC	CTACTGTGAA	ATCATGTCTG	TATTTTTCTG	1140
10	GAGCAGTTCA	CATAGAGTAG	AATGTGGAAT	TTCCCGTGAA	CGTCTCCTTC	CTCCCCCGTA	1200
	TCTGCCGCCT	GTCACTTCGC	CACCGTGCTA	GAATACTGTT	GTGTTGTAAG	ATGACTAATT	, 1260
15	TTAAAAGAAC	CTGCCCTGAA	AAGTTCTTAG	AAACGCAATG	AAAGGGAGGA	ACTTGTCCTT	1320
	TACCCAGTTT	TTCCTTTGTA	GGATGGGAAA	GTATAAAAAG	GCACAGAAGG	TTGTCATGGG	1380
20	CTGTTCCTTG	GGGGTTTTTA	TCCTGCTCAC	CGTGGAGATA	AGCCTGCGGC	TTGTCTAACC	1440
20	AGCGĆAGCGM	AAAGGTCTCA	ATGCCTTTTG	GTAACATCCG	TCATTGCAGA	AGAAAGTTTA	1500
	CACGACGTCA	AAAAGTGACG	TTCATGCTAA	GTGTTTTTCC	AGAAATATTG	GTTTCATGTT	1560
25	TCTTATTKGC	TCTGCCTCCT	GTGCTTATAT	CATCCAAAAA	CTTTTTAAAA	AGGTCCAGAA	1620
	TTCTATTTTA	ACCTGATGTT	GAGCACCTTT	' AAAACGTTCG	TATGTGTGTI	GCACTAATTC	1680
30	TAAACTTTGG	AGGCATTITG	CTGTGTGAGG	CCGATCGCCA	. CTGTAAAGGI	CCTAGAGTTG	1740
30	CCTGTTTGTC	TCTGGAGATG	GAATTAAACC	AAATAAAGAG	CTTCCACTGG	AGGCTTGTAT	1800
	TGACCTTGTA	ACTATATGTT	AATCTCGTGT	TAAAATAAAA	TATAACTTG1	GAAAAAAAA	1860
35	AAAAAAAA	NT					1872

40 (2) INFORMATION FOR SEQ ID NO: 23:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 289 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 23:

60

(2) INFORMATION FOR SEQ ID NO: 24:

5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 3533 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 24:	
	TTTTATTTAC TTCAAATTAA CTGTACTTTA CTCAAATAGA AAANGAATAA TTTTCACATT	60
15	ATGAAGCTAC ACAATTCCAA AATACACATG CTGAGGCTCT TTTTAAGTCC GAATTGTCTA	120
	GTAATTACAA AAAAGTGAAG AGTTTACAGA TATACAAGGA AATAAAGGCG AATTATTGCA	180
20	AAGAAAACAA GTTTAATTTC ACTTTGAATG ACAACGATTT TTCTGGAAAG CAGATACTTC	240
	ACTCCTTTAA GTTTCCACCC AAGCCACAAT AATTTCAAAC GGTCTTGCGG ATGACCCAGC	300
	TGGTCACTCT TGTTTATGTG GGGACTGGAG GTAATGAGAG CCAAAAAAAG TGCTATAAAC	360
25	CTAATTTGGC TAGAGCAAGT TCACACGACA CGACCGTGCT TTAAAAACTT GCTCTCCATT	420
	ATGTACTTCC TTCCATCAGG TTGGGGAAAA AAAAATGGTG GGGATGGTGA GTAAACACAC	480
30	CAGTGGTTTC ATCAGAGGG AACTCACTAC TCAGGAGGTG ACGGTGACGT GGTGCCGGTC	540
	CCTGAAGTAC GCGCACAAGC TCCGGAGGTT GCGGGAGGTT CCGCTGCCGC CTGGAGGGAA	600
	GCCGGAGCGA CGGGGGTCAC GGCGGCGGTC AGAGGGTAAA GGTCTTGCTC CCAGCAGCCT	660
35	CCGCGGTGGA TACGTCGCCA TCTTGGATCC GCGGGACAAG AAAATTCATG CGAGGGAGAC	720
	GTGGTGGGCG GTCCTTCCTG TGACACGACC CTTGAGTGAC AGTTCTATTT GATTGCCTCC	780
40	GGTACTGTGA GGAAAGGACA CGACTCTATG GTGAGGACTG ATGGACATAC ATTATCTGAG	840
	AAAAGAAACT ACCAGGTGAC AAACAGCATG TTTGGTGCTT CAAGAAAGAA GTTTGTAGAG	900
	GGGGTCGACA GTGACTACCA TGACGAAAAC ATGTACTACA GCCAGTCTTC TATGTTTCCA	960
45	CATCGGTCAG AAAAAGATAT GCTGGCATCA CCATCTACAT CAGGTCAGCT GTCTCAGTTT	1020
	GGGGCAAGTT TATACGGGCA ACAAAGTGCA CTAGGCCTTC CAATGAGGGG GATGAGCAAC	1080
50	AATACCCCTC AGTTAAATCG CAGCTTATCA CAAGGCACTC AGTTACCGAG CCACGTCACG	1140
	CCAACAACAG GGGTACCAAC AATGTCACTT CACACGCCTC CATCTCCAAG CAGGGGTATT	1200
	TTGCCTATGA ATCCTARGAA TATGATGAAC CACTCCCAGG TTGGTCAGGG CATTGGAATT	1260
55	CCTAGCAGGA CAAATAGCAT GAGCAGTTCA GGGTTAGGTA GCCCCAACAG AAGCTCGCCA	1320
	AGCATAATAT GTATGCCAAA GCAGCAGCCT TCTCGACAGC CTTTTACTGT GAACAGTATG	1380
60	TCTGGATTTG GAATGAACAG GAATCAGGCA TTTGGAATGA ATAACTCCTT ATCAAGTAAC	1440

	ATTYTTAATG GAACAGACGG AAGTGAAAAT GTGACAGGAT TGGACCTTTC AGATTTCCCA	1500
	GCATTAGCAG ACCGAAACAG GAGGGAAGGA AGTGGTAACC CAACTCCATT AATAAACCCC	1560
5	TTGGCTGGAA GAGCTCCTTA TGTTGGAATG GTAACAAAAC CAGCAAATGA ACAATCCCAG	1620
	GACTTCTCAA TACACAATGA AGATTTTCCA GCATTACCAG GCTCCAGCTA TAAAGATCCA	1680
10	ACATCAAGTA ATGATGACAG TAAATCTAAT TTGAATACAT CTGGCAAGAC AACTTCAAGT	1740
10	ACAGATGGAC CCAAATTCCC TGGAGATAAA AGTTCAACAA CACAAAATAA TAACCAGCAG	1800
	AAAAAAGGGA TCCAGGTGTT ACCTGATGGT CGGGTTACTA ACATTCCTCA AGGGATGGTG	1860
15	ACGGACCAAT TTGGAATGAT TGGCCTGTTA ACATTTATCA GGGCAGCAGA GACAGACCCA	1920
	GGAATGGTAC ATCTTGCATT AGGAAGTGAC TTAACAACAT TAGGCCTCAA TCTGAACTCT	1980
20	CCTGAAAATC TCTACCCCAA ATTTGCGTCA CCCTGGGCAT CTTCACCTTG TCGACCTCAA	2040
20	GACATAGACT TCCATGTTCC ATCTGAGTAC TTAACGAACA TTCACATTAG GGATAAGCTG	2100
	GCTGCAATAA AACTTGGCCG ATATGGTGAA GACCTTCTCT TCTATCTCTA TTACATGAAT	2160
25	GGAGGAGACG TATTACAACT TTTAGCTGCA GTGGAGCTTT TTAACCGTGA TTGGAGATAC	2220
	CACAAAGAAG AACGAGTATG GATTACCAGG GCACCAGGCA TGGAGCCAAC AATGAAAAACC	2280
30	AATACCTATG AGAGGGGAAC ATATTACTTC TTTGACTGTC TTAACTGGAG GAAAGTAGCT	2340
50	AAGGAGTTCC ATCTGGAATA TGACAAATTA GAAGAACGGC CTCACCTGCC ATCCACCTTC	2400
	AACTACAACC CTGCTCAGCA AGCCTTCTAA AAAAAAAAAA	2460
35	CCCTTTTCTT GGGGTATGGC TGTCTCAGCA CAATACTCAA CATAACTGCA GAACTGATGT	2520
	GGCTCAGGCA CCCTGGTTTT AATTCCTTGA GGATCTGGCA ATTGGCTTAC GCAAAAGGTC	2580
40	ACCATTIGAG GTCCTGCCTT ACTAATTATG TGCTGCCCAA CAACTAAATT TGTAATTIGT	2640
40	TTTTCTCTAG TTTGAGCAGG GTCTGAATTT TTTCATTTAT TTCCTTTTTT GCCAGCAGAC	2700
	AGACTIGAGT CTGTAAAGAC AAGCAAATAC ACTGACAGAA GTTTACCATA GTTTCTAAAA	2760
45	TGTAAAAAG AAAACCCCCA AAAGACTCAA GAAAATTAGA CCACAAATTT TGCATTGTTC	2820
	ATTGTAGCAC TATTGGTAAT AAAATÄACAA ATGTTTGTGC ATTTTTATGT GAAGATCCTT	2880
50	CTCGTATTTC ATTTGGAAAG ATGAGCAAGA GGTCTGCTTC CTTCATTTTA CTTCCCCTTC	2940
50	TGTTTTGAA AGGCAGTTTC GCCAAGCTTA ATGCAAGAAT ATCTGACTGT TTAGAAGAAA	3000
	GATATTGCCA CAATCTCTGG ATGGTTTTCC AGGGTTGTGT TATTACTGAG CTTCATCTTT	3060
55	CCAGAATGAG CAAAACACTG TCCAGTCTTT GTTACGATTT TGTAATAAAT GTGTACATTT	3120
	TITITTAAATT TTTGGACATC ACATGAATAA AGGTATGTAT GTACGAATGT GTATATATTA	3180
60	TATATATGAC ATCTATTTTG GAAAATGTTT GCCCTGCTGT ACCTCATTTT TAGGAGGTGT	3240

	GCATGGATGC AATATATGAA AATGGGACAT TCTGGAACTG CTGGTCAGGG GACTTTGTCG	3300
	CCCTGTGCAC TAAAAGGGCC AGATTTTCAG CAGCCAAGGA CATCCATACC CAAGTGAATG	3360
5	TGATGGGACT TÅAAAGAAGT GAACTGAGAC AATTCACTCT GGCTGTTTGA ACAGCAGCGT	3420
	TTCATAGGAA GAGAAAAAA GATCAATCTT GTATTTTCTG ACCACATAAA GGCTTCTTCT	3480
10	CTTTGTAATA AAGTAGAAAA GCTCTCCTCA AAAAAAAAAA	3533
	(2) INFORMATION FOR SEQ ID NO: 25:	
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1148 base pairs (B) TYPE: nucleic acid	
20	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 25:	
25	ACCCACGCGT CCGCAAATTA TACTTCCTCA TTCATATTAT GTTGATACAA AAGACCTTGG	60
	CAGCCATTTC TCCCAGCAGT TTTAAAGGAT GAACATTGGA TTTCATGCCA TCCCATAGAA	120
	AACCTGTTTT AAAATTTTAG GGATCTTTAC TTGGTCATAC ATGAAAAGTA CACTGCTTAG	180
30	AAATTATAGA CTATTATGAT CTGTCCACAG TGCCCATTGT CACTTCTTTG TCTCATTTCT	240
	TCCCTTTGTT CCTTAGTCAT CCAAATAAGC CTGAAAACCA TAAGAGATAT TACTTTATTG	300
35	AATATGGTTG GCATTAAATT TAGCATTTCA TTATCTAACA AAATTAATAT AAATTCCAGG	360
	ACATGGTAAA ATGTGTTTTA ATAACCCCCA GACCCAAATG AAAATTTCAA AGTCAATACC	420
	AGCAGATTCA TGAAAGTAAA TTTAGTCCTA TAATTTTCAG CTTAATTATA AACAAAGGAA	480
40	CAAATAAGTG GAAGGGCAGC TATTACCATT CGCTTAGTCA AAACATTCGG TTACTGCCCT	540
	TTAATACACT CCTATCATCA GCACTTCCAC CATGTATTAC AAGTCTTGAC CCATCCCTGT	600
45	CGTAACTCCA GTAAAAGTTA CTGTTACTAG AAAATTITTA TCAATTAACT GACAAATAGT	660
	TTCTTTTTAA AGTAGTTTCT TCCATCTTTA TTCTGACTAG CTTCCAAAAT GTGTTCCCTT	720
	TTTGAATCGA GGTTTTTTTG TTTTGTTTTG TTTTCTGAAA AAATCATACA ACTTTGTGCT	780
50	TCTATTGCTT TTTTGTGTTT TGTTAAGCAT GTCCCTTGGC CCAAATGGAA GAGGAAATGT	840
	TTAATTAATG CTTTTTAGTT TAAATAAATT GAATCATTTA TAATAATCAG TGTTAACAAT	900
55	TTAGTGACCC TTGGTAGGTT AAAGGTTGCA TTATTTATAC TTGAGATTTT TTTCCCCTAA	960
	CTATTCTGTT TTTTGTACTT TAAAACTATG GGGGAAATAT CACTGGTCTG TCAAGAAACA	1020
	GCAGTAATTA TTACTGAGTT AAATTGAAAA GTCCAGTGGA CCAGGCATTT CTTATATAAA	1080
60	TAAAATTIGT GGTACTAATG TGAAAAAAA AAAAAAAAA AACTGGAGGG GGGGGGGG	1140

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	CCCTATTA	1148
5		
	(2) INFORMATION FOR SEQ ID NO: 26:	
10	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 717 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 26:	
	GGCACGAGCT AGCTGCCGCC ACCCGAACAG CCTGTCCTGG TGCCCCGGCT CCCTGCCCCG	60
20	CGCCCAGTCA TGACCCTGCG CCCCTCACTC CTCCCGCTCC ATCTGCTGCT GCTGCTGCTG	120
20	CTCAGTGCGG CGGTGTGCCG GGCTGAGGCT GGGCTCGAAA CCGAAAGTCC CGTCCGGACC	180
	CTCCAAGTGG AGACCCTGGT GGAGCCCCCA GAACCATGTG CCGAGCCCGC TGCTTTTGGA	240
25	GACACGCTTC ACATACACTA CACGGGAAGC TTGGTAGATG GACGTATTAT TGACACCTCC	300
	CTGACCAGAG ACCCTCTGGT TATAGAACTT GGCCAAAAGC AGGTGATTCC AGGTCTGGAG	360
30	CAGAGTCTTC TCGACATGTG TGTGGGAGAG AAGCGAAGGG CAATCATTCC TTCTCACTTG	420
30	GCCTATGGAA AACGGGGATT TCCACCATCT GTCCCAGCGG ATGCAGTGGT GCAGTATGAC	480
	GTGGAGCTGA TTGCACTAAT CCGAGCCAAC TACTGGCTAA AGCTGGTGAA GGGCATTTTG	540
35	CCTCTGGTAG GGATGGCCAT GGTGCCAGCC CTCCTGGGCC TCATTGGGTA TCACCTATAC	600
	AGAAAGGCCA ATAGACCCAA AGTCTCCAAA AAGAAGCTCA AGGAAGAGAA ACGAAACAAG	660
40	AGCAAAAAGA AATAATAAAT AATAAATTTT AAAAAAAAA AAAAAA	717
45	(2) INFORMATION FOR SEQ ID NO: 27: (i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 1099 base pairs (B) TYPE: nucleic acid	
50	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 27:	
	GGCACGAGCC GATGTGGACA TCATCCTGTC TATCCCCATG TTCCTGCGCC TGTACCTGAT	60
55	CGCCCGAGTC ATGCTGCTGC ACAGCAAGCT CTTCACCGAT GCCTCGTCCC GCAGCATCGG	120
	GGCCCTCAAC AAGATCAACT TCAACACCCG CTTTGTCATG AAGACGCTCA TGACCATCTG	180
60	CCCTGGCACT GTGCTGCTCG TGTTCAGCAT CTCTCTGTGG ATCATTGCTG CCTGGACCGT	240

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	CCGTGTCTGT	GAAAGTCCTG	AATCACCAGC	CCAGCCTTCT	GGCTCATCAC	TTCCTGCTTG	300
5	GTACCATGAC	CAGCAGGACG	TAACTAGTAA	CTTTCTGGGT	GCCATGTGGC	TCATCTCCAT	360
	CACATTCCTT	TCCATTGGTT	ATGGGGACAT	GGTGCCCCAC	ACATACTGTG	GGAAAGGTGT	420
	CTGTCTCCTC	ACTGGCATCA	TGGGTGCAGG	CTGCACTGCC	CTTCTGGTGG	CCGTGGTGGC	480
10	CCGAAAGCTG	GAACTCACCA	AAGCGGAGAA	GCACGTTCAT	AACTTCATGA	TGGACACTCA	540
	GCTCACCAAG	CGGATCAAGA	ATGCTGCAGC	CAATGTCCTT	CGGGAAACAT	GGTTAATCTA	600
15	TAAACACACA	AAGCTGCTAA	AGAAGATTGA	CCATGCCAAA	GTGAGGAAAC	ACCAGAGGAA	660
	GTTCCTCCCA	AGCTATCCAC	CAGTTTGAGG	AGCGTCCCAG	ATGGAACAGA	GGAAAGCTGA	720
	GTGACCAAGC	CAACACTCTG	GTGGACCTTT	CCAAGATGCA	GAATGTCATG	TATGACTTAA	780
20	TCACAGAACT	CAATGACCGG	AGCGAAGACC	TGGAGAAGCA	GATTGGCAGC	CTGGAGTCGA	840
	AGCTGGAGCA	TCTCACCGCC	AGCTTCAACT	CCCTGCCGCT	GCTCATCGCC	GACACCCTGC	900
25	GCCAGCAGCA	GCAGCAGCTC	CTGTCTGCCA	TCATCGAGGC	CCGGGGTGTC	AGCGTGGCAG	960
	TGGGCACCAC	CCACACCCCA	ATCTCCGATA	GCCCCATTGG	GGTCAGCTCC	ACCTCCTTCC	1020
	CGACCCCGTN	CACAAGTTCA	AGCAGTTGCT	AAATAAATCT	CCCCACTCCA	GAAGCATTAA	1080
80	AAAAAAAA	AAAAAAAA					1099

35 (2) INFORMATION FOR SEQ ID NO: 28:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 941 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 28:

45	AATTCGGCAG	AGAGCCAACC	GAGGGCGTTC	CTGTCGGGGC	TGCAGCGGCG	GGAGGGAGCC	60
	CAGTGGAGGC	GCCCTCCCGA	AGCGCCACTG	CCCATGCTGA	CCACCCAGCC	CTCCGGCTGC	120
50	TGATGTCATG	AGTAACACCA	CTGTGCCCAA	TGCCCCCAG	GCCAACAGCG	ACTCCATGGT	180
	GGGCTATGTG	TTGGGGCCCT	TCTTCCTCAT	CACCCTGGTC	GGGTGGTGG	TGGCTGTGGT	240
	AATGTATGTA	CAGAAGAAAA	AGCGGGTGGA	CCGGCTGCGC	CATCACCTGC	TCCCCATGTA	300
55	CAGCTATGAC	CCAGCTGAGG	AACTGCATGA	GGCTGAGCAG	GAGCTGCTCT	CTGACATGGG	360
	AGACCCCAAG	GTGGTACATG	GCTGGCAGAG	TGGCTACCAG	CACAAGCGGA	TGCCACTGCT	420
60	GGATGTCAAG	ACGTGACCTG	ACCCCCTTGC	CCCACCCTTC	AGAGCCTGGG	GTYCTGGACT	480

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	GCCTGGGGCC	CTGCCATCTG	CTTCCCCTGC	TGTCACCTGG	STCCCCCTGC	TGGGTGCTGG	540
	GTCTCCATTT	CTCCCTCCAC	CCACCCTCAG	CAGCATCTGC	TTCCCATGCC	CTCACCATCA	600
5	CCTCACTGCC	CCCAGGCCTT	CTGCCCTTTG	TGGGTGTTGA	GCTCACCGCC	CACCCACAGG	660
	CACTCATGGG	AAGAGGCTTT	CCTTCTGGGA	TGGCGGCGGC	TGGTAGACAC	CTTTGCTTTC	720
10	TCTAGCCCTC	CTGGGCTGGG	CTTGGGCACA	AATCCCCAGG	CAGGCTTTGG	AGTTGTTTCC	780
10	ATGGTGATGG	GGCCAGATGT	ATAGTATTCA	GTATATATTT	TGTAAATAAA	ATGTTTGTG	840
	GCTAAAAAAA	ААААААААА	ATCNAAGGGG	GGGCCGGTAC	CCAAATTCCC	CCTATANTGA	900
15	ATTCGTATTA	ACAATTCACT	TGGGGCCGTC	CTTTTAANAA	С		941

20 (2) INFORMATION FOR SEQ ID NO: 29:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 756 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 29:

30	GGCACGAGGA AGCTGGAGCG GGCCGGCGGT GCAGTCACGG GGGAGCGAGG CCTGCTGGGC	60
	TTGGCAACGA GGGACTCGGC CTCGGAGGCG ACCCAGACCA CACAGACACT GGGTCAAGGA	120
35	GTAAGCAGAG GATAAACAAC TGGAAGGAGA GCAAGCACAA AGTCATCATG GCTTCAGCGT	180
33	CTGCTCGTGG AAACCAAGAT AAAGATGCCC ATTTTCCACC ACCAAGCAAG CAGAGCCTGT	240
	TGTTTTGTCC AAAATCAAAA CTGCACATCC ACAGAGCAGA GATCTCAAAG ATTATGCGAG	300
40	AATGTCAGGA AGAAAGTTTC TGGAAGAGAG CTCTGCCTTT TTCTCTTGTA AGCATGCTTG	360
	TCACCCAGGG ACTAGTCTAC CAAGGTTATT TGGCAGCTAA TTCTAGATTT GGATCATTGC	420
15	CCAAAGTTGC ACTTGCTGGT CTCTTGGGAT TTGGCCTTGG AAAGGTATCA TACATAGGAG	480
45	TATGCCAGAG TAAATTCCAT TTTTTTGAAG ATCAGCTCCG TGGGGCTGGT TTTGGTCCAC	540
	AGCATAACAG GCACTGCCTC CTTACCTGTG AGGAATGCAA AATAAAGCAT GGATTAAGTG	600
50	AGAAGGGAGA CTCTCAGCCT TCAGCTTCCT AAATTCTGTG TCTGTGACTT TCGAAGTTTT	660
	TTAAACCTCT GAATTTGTAC ACATTTAAAA TTTCAAGTGT ACTTTAAAAT AAAATACTTC	720
55	ТААТОGAAAA AAAAAAAAAAAAAAAAAAAAAAAAAAAAAAAA	756

(2) INFORMATION FOR SEQ ID NO: 30:

60

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2100 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 30:

10	NCCAGAGGCA	GAAAGTCCTG	CTTCTGGGGC	GTAACCTACA	GGATATCCTT	GGAACAGAAG	60
10	ATCTTATTGT	GGAAGTRACT	TCCAATGATG	CTGTGAGATT	TTATCCCTGG	ACCATTGATA	120
	ATAAATACTA	TTCAGCAGAC	ATCAATCTAT	GTGTGGTGCC	AAACAAATTT	CTTGTTACTG	180
15	CAGAGATTGC	AGAATCTGTC	CAAGCATTTG	TGGTTTACTT	TGACAGCACA	CAAAAATCGG	240
	GCCTTGATAG	TGTCTCCTCA	TGGCTTCCAC	TGGCAAAAGC	ATGGTTACCY	GAGGTGATGA	300
20	TCTTGGTCTG	CGATAGAGTG	TCTGAAGATG	GTATAAACCG	ACAAAAAGCT	CAAGAATGGT	360
	GCATCCAAAC	ATGGCTTTGA	ATTGGTAGAA	CTTAGTCCAG	AGGAGTTGCC	TGAGGAGGAT	420
	GATGACTTCC	CAGAATCTAC	AGGAGTAAAG	CGAATTGTCC	AAGCCCTGAA	TGCCAATGTG	480
25	TGGTCCAATG	TAGTGATGAA	GAATGATAGG	AACCAAGGCT	TTAGCTTGCT	GCAACTCATT	540
	GACTGGAACA	AACCATAGCA	TTGGGTCAGC	AGATCCCTGT	CACCCAGAGC	AACCCCATTT	600
30	GCCAGCAGCA	GATAGTACTG	AATCCCTCTC	TGATCATCGG	GGTGGTGCAT	CTAACACAAC	660
	AGATGCCCAG	GTTGATAGCA	TTGTGGATCC	CATGTTAGAT	CTGGATATTC	AAGAATTAGC	7 20
	CAGTCTTACC	ACTGGAGGAG	GAGATGTGGA	GAATTTTGAA	AGACTCTTTT	CAAAGTTAAA	780
35	GCAAATGAAA	GACAAGGCTG	CGACGCTTCC	TCATGAGCAA	AGAAAAGTGC	ATGCAGAAAA	840
	GGTGGCCAAA	GCATTCTGGA	TGGCAATCGG	GGGAGACAGA	GATGAAATTG	AAGGCCTTTC	900
40	ATCTGATGAA	GAGCACTGAA	TTATTCATAC	TAGGGTTTGA	CCAACAAAGA	TGCTAGCTGT	960
	CTCTGAGATA	CCTCTCTACT	CAGCCCAGTC	ATATTTTGCC	AAAATTGCCC	TTATCATGTT	1020
	GGCTGCCTGA	CTTGTTTATA	GGGTCCCCTT	AATTTTAGTT	TTTAGTAGGA	GGTTAAGGAG	1080
45	AAATCTTTTT	TTTCCTCAGT	ATATTGTAAG	AGAGTGAGGA	ATACAGTGAT	AGTAATGAGT	1140
	GAGGATTTCT	TAAATRTACT	TTTTTTTTGT	TCTAGGAATG	AGGGTAGGAT	AAATCTCAGA	1200
50	GGTCTGTGTG	ATTTACTCAA	GTTGAAGACA	ACCTCCAGGC	CATTCCTGGT	CAACCTTTTA	1260
	AGTAGCATTT	CCAGCATTCA	CACTTGATAC	TGCACATCAG	GAGTTGTGTC	ACCTTTCCTG	1320
	GGTGATTTGG	GTTTTCTCCA	TTCAAGGAGC	TTGTAGCTCT	GAAGCTATGA	TGCTTTTATT	1380
55	GGGAGGAAAG	GAGGCAGCTG	CAGAATTGAT	GTGAGCTATG	TGGGGCCGAA	GTCTCAGCCC	1440
	GCAGCTAAGT	CTCTACCTAA	GAAAATGCCT	CTGGGCATTC	TTTTGAAGTA	TAGTGTCTGA	1500
60	GCTCATGCTA	GAAAGAATCA	AAAAGCCAGT	GTGGATTTTT	AGACTGTAAT	AAATGAGGCA	1560

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	AAGGATTTCT	ATTCCAGTGG	GAAGRAAACC	TCTCTACTGA	GTTGTGGGGG	ATATGTTGTA	1620
	TGTTAGAGAG	AACCTTAAGG	AGTCCTTGTA	TGGGCCATGG	AGACAGTATG	TGATAACATA	1680
5	CCGTGATTTT	CATGAAGAAA	TTCTTCTGTC	TTAGAGTTCT	CCCCTGCTGC	TTGAGATGCC	1740
	AGAGCTGTGT	TGTTGCACAC	CTGCAAAACA	AGGCACATTT	CCCCCTTTCT	CTTTAAAGCC	1800
10	AAAGAGAGAT	CACTGCCAAA	GTGGGAGCAC	TAAGGGTGG	GTGGGGAAGT	GAAATGTTAG	1860
10	GCGATGAATT	CCTGAGCACC	TIGTTTTTCT	TCCAAGGTTC	GTAGCTCCTC	TCTGCCCTTC	1920
	CAAGCCTGTA	ACCTCGGAGG	ACTATCTTTT	GTTCTTTATC	CTTTGTCTTG	TTTGAGTGGG	1980
15	TCAGCCCCAG	AGGAACTGAT	AAGCAAATGG	CAAGTTTTTA	AAGGAAGAGT	GGAAAGTACT	2040
	GCAAATAAAA	ATCCTTATIT	GTTTTTGTAG	ааааааааа	AAAAAAAAA	AAAAAAAAG	2100

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(2) INFORMATION FOR SEQ ID NO: 31:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1448 base pairs

(B) TYPE: nucleic acid(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

30 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 31:

	ААААААААА	AAAGCCCACC	TGAAAGCCTG	TCTCTTTCCA	CTTTGTTGGC	CCTTCCAGTG	60
35	GGATTATCGA	GCATGTTGTT	TTTTCATAGT	GCCTTTTTCC	TTATTTCAAG	GGTTGCTTCT	120
33	GAGTGGTGTT	TATATATATATATAT	TTAATTTGTT	TTGTTTTAAA	ATAAGTTAAA	GACAGTCCAG	180
	AGCTTTTCAG	CCAATTTGTC	TCCTACTCTG	TGTAAATATT	TTTCCCTCCG	GGCAGGGGAG	240
40	CCAGGGTAGA	GCAAAGGAGA	CAAGCAGGAG	TGGAAGGTGA	GGCGTTCTCC	TGCTTGTACT	300
	AAGCCAGGAG	STTTAAGCTC	CAGCTTTAAG	GGTTGTGAGC	CCCTTGGGGT	TCAGGGAACT	360
45	GCTTGCCCAG	GGTGCAGTGT	GAGTGTGATG	GGCCACCGGG	GCAAGAGGGA	AGGTGACCGC	420
73	CCAGCTCTCC	CACATCCCAC	TGGATCTGGC	TTACAGGGG	GTCGGAAGCC	TGTCCTCACC	480
	GTCTCGGGG	TTGTGGCCCC	CGCCCCTCC	CTATATGCAC	CCCTGGAACC	AGCAAGTCCC	540
50	AGACAAGGAG	AGCGGAGGAG	GAAGTCATGG	GAACGCAGCC	TCCAGTTGTA	GCAGGTTTCA	600
	CTATTCCTAT	GCTGGGGTAC	ACAGTGAGAG	TACTCACTTT	TCACTTGTCT	TGCTCTTAGA	660
55	TTGGGCCATG	GCTTTCATCC	TGTGTCCCCT	GACCTGTCCA	GGTGAGTGTG	AGGGCAGCAC	720
33	TGGGAAGCTC	GAGTGCTGCT	TGTGCCTCCC	TTCCCAGTGG	GCTGTGTTGA	CTGCTGCTCC	780
	CCACCCCTAC	CGATGGTCCC	AGGAAGCAGG	GAGAGTTGGG	GAAGGCAAGA	TTGGAAAGAC	840
60	AGGAAGACCA	AGGCCTCGGC	AGAACTCTCT	GICTICICIC	CACTTCTGGT	CCCCTGTGGT	900

	GATGTGCCTG TAATCTTTTT CTCCACCCAA ACCCCTTCCC ACGACAAAAA CAAGACTGCC	960
_	TCCCTCTCTT CCGGGAGCTG GTGACAGCCT TGGGCCTTTC AGTCCCAAAG CGGCCGATGG	1020
5	GAGTCTCCCT CCGACTCCAG ATATGAACAG GGCCCAGGCC TGGAGCGTTT GCTGTGCCAG	1080
	GAGGCGGCAG CTCTTCTGGG CAGAGCCTGT CCCCGCCTTC CCTCACTCTT CCTCATCCTG	1140
10	CTTCTCTTTT CCTCGCAGAT GATAAAAGGA ATCTGGCATT CTACACCTGG ACCATTTGAT	1200
	TGTTTTATTT TGGAATTGGT GTATATCATG AAGCCTTGCT GAACTAAGTT TTGTGTGTAT	1260
	ATATTTAAAA AAAAAATCAG TGTTTAAATA AAGACCTATG TACTTAATCC TTTAACTCTG	1320
15	CGGATAGCAT TTGGTAGGTA GTGATTAACT GTGAATAATA AATACACAAT GAATTCTTMA	1380
	AAAAAAAAA AAAAAAAAA AAAAAAAAA AAACCCCGGG GGGGCCCCCG GGCCCCAATT	1440
20	CCCCCCAA	1448
25	(2) INFORMATION FOR SEQ ID NO: 32:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 456 base pairs	
30	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 32:	
35	GGCACAGCAA ACTTGACGCC ATGAAGATCC CGGTCCTTCC TGCCGTGGTG CTCCTCTCCC	60
	TCCTGGTGCT CCACTCTGCC CAGGGAGCCA CCCTGGGTGG TCCTGAGGAA GAAAGCACCA	120
40	TTGAGAATTA TGCGTCACGA CCCGAGGCCT TTAACACCCC GTTCCTGAAC ATCGACAAAT	180
	TGCGATCTGC GTTTAAGGCT GATGAGTTCC TGAACTGGCA CGCCCTCTTT GAGTCTATCA	240
	AAAGGAAACT TCCTTTCCTC AACTGGGATG CCTTTCCTAA GCTGAAAGGA CTGAGGAGCG	300
45	CAACTCCTGA TGCCCAGTGA CCATGACCTC CACTGGAAGA GGGGGCTAGC GTGAGCGCTG	360
	ATTCTCAACC TACCATAACT CTTTCCTGCC TCAGGAACTC CAATAAAACA TTTTCCATCC	420
50	AAAAAAAAA AAAAAAAAAC CCCNGGGGGG GCCCGG	456
	(2) INTODIATION FOR STO TO NO. 32	
55	(2) INFORMATION FOR SEQ ID NO: 33:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1326 base pairs	
	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
50	(D) TOPOLOGY: linear	

	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 33:	
~	GGCACGAGTG CAGGCCCAGA GAGGACTCAT TGAAAGGACT GAAAGGGGAG GTGGCGTTTT	60
5	CTTCCTACCC AAACTTACCC CTGTGAGCTG GACAGCTTGG TAGCACCTGC CTGGACTTAG	120
	ATGGTGGTAG CCAAGAAGAC TGACATTTTA GGGAACAGGA CGGGGAGGAG AAGGCTCTGG	180
10	CACACACACA TGTGTCCATA TGTCCTGCAA TGGTCTGGGG ACTATTGCTA GGCTAGGAGC	240
	CCTAAGTGTC TTCTTCCTCA TGTCTMTTCT CCCCTGTSTC ATGGGCCCTA AGRTCTCTTT	300
1.5	CACTGGGCCT GCCTCAATGA ACGTGCTGCC CAGCTACCCC GAAACACGGC ANCTGCCGGC	360
15	TATCAATGCC CCAGCTGCAA TGGCCCATCT TCCCCCAACC AACCTGGCTG GGCCCGTGGG	420
	CTCCGCACTG AGARARAAAS TTGGCACART CAACTGGGCC CGGGCAGGAC TGGGCCYCCC	480
20	TCTGATCGAT GAAGKTGGTG ARCCCAGAGC CCGAGCCCCT CAACACGTCT GACTTCTCTG	540
	ACTGGTCTAG TTTTAATGCC AGCAGTACCC CTGGACCAGA GGAGGTAGAC AGCGCCTCTG	600
25	CTGCCCCAGC CTTCTACAGC CGAGCCCCCC GGCCCCAGC TTCCCCAGGC CGGCCCGAGC	660
23	AGCACACAGT GATCCACATG GGCAATCCTG AGCCCTTGAC TCACGCCCCT AGGAAGGTGT	720
	ATGATACGCG GGATGATGAC CGGACACCAG GCCTCCATGG AGACTGTGAC GATGACAAGT	780
30	ACCGACGTCG GCCGGCCTTG GGTTGGCTGG CCCGGCTGCT AAGGAGCCGG GCTGGGTCTC	840
	GGAAGCGRCC GCTGACCCTG CTCCAGCGGG CGGGGCTGCT GCTACTCTTG GGACTGCTGG	900
35	GCTTCCTGGC CCTCCTTGCC CTCATGTCTC GCCTAGGCCG GGCCGCAGCT GACAGCGATC	960
33	CCAACCTGGA CCCACTCATG AACCCTCACA TCCGCGTGGG CCCCTCCTGA GCCCCCTTGC	1020
	TTGTGGCTAG GCCAGCCTAG GATGTGGGTT CTGTGGAGGA GAGGCGGGGT AATGGGGAGG	1080
40	CTGAGGGCAC CTCTTCACTG CCCCTCTCCC TCAAGCCTAA GACACTAAGA CCCCAGACCC	1140
	AAAGCCAAGT CCACCAGAGT GGCTGCAGGC CAGGCCTGGA GTCCCCGTGG GTCAAGCATT	1200
	TOTOTTGACT TECTTTCCTC CCGGGTYTCC AGCCTCCGAC CCCTCGCCCC ATGAAGGAGC	1260

AAANAA

45

(2) INFORMATION FOR SEQ ID NO: 34:

55 (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 710 base pairs

1320

1326

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 34:	
	GCGAAAGAGA AAAAGGCTGG AGCTCCCGCC CCCGGGGCTG TCAGATGGCT TGGGTTTCTG	60
5	CGACGCGATT GGCTCGCGGA GGGCAGAAAT TACTCAGCAA ACATGACTAT TATTAGCTGC	120
	TTAGCAACAG CTCACCAAAG TAGAGAGACC ACCCAGGTAG GCAACCCAGT GTGTGCATCC	180
10	TCGGCTTCGG GGCAGCCTCT GAGAGCGCCA ACCTTCTCGC ATGCAATACT TCCATTAAGG	240
10	AATGCTCCCC CTCCTTTCTC TCTTATTCCT TTTCTTTTCA ACAGTGTCTT CTTTTTGTGG	300
	GATGCCTTTG CGCGCACACA CGCGCGCGCA SGCACACACA CGAACATTTG CCTCGCGGTA	360
15	GACACGGGGG GAAATGTWAT ATTTTTTTAA GCGCTTAAAC AATTTCTGAA ATTCCTCAAA	420
	GAAAAGCCTT TCAGARGCAC CTTGGCCTCA AGCTGCAACA AATACTGGGA RGTCCGGCTC	480
20	GCATTCCCAG GCCTGCACCA ATAATGACAG CGTGCTGGAT ARTGCGCCAG TGTGTGCCAG	540
	ATTTTTTTT CCTCTTCTCT TTTCTTTTAT AACTAAAGGG AAGACTTAGG CTCTTGCAGG	600
	GAACAACGCC TCGCATTAAG ATAAACAGAA TGGAAAGTTA AAGAGGAAAG CAAGGACGTT	660
25	GGGAAAAGCC ATCTTTCTTA AAATCCGTCT GCCCCCCAGC CGCTTTCTCC	710
30	(2) INFORMATION FOR SEQ ID NO: 35:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 1188 base pairs (B) TYPE: nucleic acid	
35	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 35:	
10	GATGGCTTTT ATATCTATTA TCGACCCACA GACAGTGACA ATGATAGTGA CTACAAGAAG	60
	GATATGGT®G AAGGGGACAA GTACTGGCAC TCCATCAGCC ACCTGCAGCC AGAGACCTCC	120
	TACGACATTA AGATGCAGTG CTTCAATGAA GGAGGGGAGA GCGAGTTCAG CAACGTGATG	180
1 5	ATCTGTGAGA CCAAAGCTCG GAAGTCTTCT GGCCAGCCTG GTCGACTGCC ACCCCCAACT	240
	CTGGCCCCAC CACAGCCGCC CCTTCCTGAA ACCATAGAGC GGCCGGTGGG CACTGGGGCC	300
50	ATGGTGGCTC GCTCCAGCGA CCTGCCCTAT CTGATTGTCG GGGTCGTCCT GGGCTCCATC	360
	GTTCTCATCA TCGTCACCTT CATCCCCTTC TGCTTGTGGA GGGCCTGGTC TAAGCAAAAA	420
	CATACAACAG ACCTGGGTIT TCCTCGAAGT GCCCTTCCAC CCTCCTGCCC GTATACTATG	480
55	GTGCCATTGG GAGGACTCCC AGGCCACCAG GCAGTGGACA GCCCTACCTC AGTGGCATCA	540
	GTGGACGGGC CTGTGCTAAT GGGATCCACA TGAATAGGGG CTGCCCCTCG GCTGCAGTGG	600
50	GCTACCCGGG CATGAAGCCC CAGCAGCACT GCCCAGGCGA GCTTCAGCAG CAGAGTGACA	660

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	CCAGCAGCCT	GCTGAGGCAG	ACCCATCTTG	GCAATGGATA	TGACCCCCAA	AGTCACCAGA	720
5	TCACGAGGGG	TCCCAAGTCT	AGCCCGGACG	AGGGCTCTTT	CTTATACACA	CTGCCCGACG	780
	ACTCCACTCA	CCAGCTGCTG	CAGCCCCATC	ACGACTGCTG	CCAACGCCAG	GAGCAGCCTG	840
	CTGSTGTGGG	CCAGTCAGGG	GTGAGGAGAG	CCCCGACAG	TCCTGTCCTG	GAAGCAGTGT	900
0	GGGACCCTCC	ATTTCACTCA	GGGCCCCCAT	GCTGCTTGGG	CCTTGTGCCA	GTTGAAGAGG	960
	TGGACAGTCC	TGACTCCTGC	CAAGTGAGTG	GAGGAGACTG	GTGTCCCCAG	CACCCCGTAG	1020
15	GGGCCTACGT	AGGACAGGAA	CCTGGAATGC	AGCTCTCCCC	GGGGCCACTG	GTGCGTGTGT	1080
13	CTTTTGAAAC	ACCACCTCTC	ACAATTTAGG	CAGAAGCTGA	TATCCCAGAA	AGACTATATA	1140
	TTGTTTTTT	TTTAAAAAAA	ААААААААА	AWCYCGGGGG	GGGGCCCC		1188

20

(2) INFORMATION FOR SEQ ID NO: 36:

25 (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 956 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

30

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36:

GGCAGAGCAG TGAAAATGCA TCCTAAAAAT TCAATGTTTA TACCAGGCTC ATGACACTAA 60 GATGTGACAT CTGGACACGA GGGGTCAGCC ACGTGGATAC ATCCCTCCCA GATTGCATCT 120 35 CCAGGAATCA CTCTGCTAGC AGAATGGGCG CCCCATCCCT TACTATGCTG CTCCTCCTCA 180 AAGTGCAGCC CAGAAGGACC CAGGCCTTTG ATGCACATTG GGTGGGTCTC CCACTACTTT 240 40 AGTTGAAATG GGAGCATGCT GGAGTCGGCG TTCTGTTGCT TCTGGTGAGA AGGACATCCC 300 ATTGACCCCT GGCCACCAGG TCCAGTATTC CATCCTTCCT TCTGTCCCAG CCTATCGCCC 360 TCCCCACYAG GCCCACCCC ACAACTTCTC CTCAAGGGAG GTTNTCCCGC AGCTGGAGGG 420 45 CTTGCACAGA CCAGCAGTCA CAGAAATCAT TCTTCCTGCT GTACTGGGCC TTAACTGCCT 480 GCAAATGTCC GAGCACTACT GCATAGGATG CCAGAGCCAC CGAAGATAAA CACAGCCAAG 540 50 TTTAATAATA ATAAAAGGAA AAATCTCAGC CTGCAGAACT CTGGTTTTGA CCCACCATCG 600 GCCAGATGCA CATCTTCAGG GCCTGTTGAG CACCTTCTGA AAAGCAGGGC TCGTAATAGA 660 CTCCAGCACA TTCCATCAGA GTCAGGAAAA CTGCGGTGAG TCCCAGAGAA TCTAGGGTGC 720 55 AGGGCAGGGA GCAGGAGTCA TAAGGAGTGA TAACCTAAAC TGTGTGTAGT CAGCGGGGAG 780 GGTCTTATGT TATCAGGTGA AATGAGAGCC AGTAAGTTAG TTGATCCTGT CACAGATATA 840 60

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	ACCCTGATAA	CACCCCATAG	ATACGCGACA	CGTGTGTCCT	GCCCCTGCTT	TCCCCATCCA	900
	ACATGGTTCT	TCTGTTCCAC	AGACATTAAA	GGGGCTTTCT	GCAATTACTT	AAAAA	956
5							
			EQ ID NO: 3				
10 15	(i)	(A) LEN (B) TYF (C) STR	HARACTERIST GTH: 1603 b E: nucleic ANDEDNESS: YOLOGY: line	ase pairs acid double			
13	(xi) SEQUENCE	DESCRIPTION	: SEQ ID NO	: 37:		
	TCGACCCACG	CGTCCGCTCT	GCCAGGAATC	TGGTCTTTCT	GTAGACCCAA	GTCAGAAAGA	60
20	ACCATTTGTG	GAGTTAAATC	GAATATTAGA	RGCATTAAAR	GTCAGAGTTC	TGAGACCTGC	120
	TCTGGAATGG	GCAGTTTCAA	ACCGAGAGAT	GCTTATAGCC	CAAAACAGCT	CCTTGGAATT	180
25	TAAACTACAC	AGACTGTATT	TTATTAGCTT	RTTAATGGGT	GGAACACAAA	TCAGCGAGAR	240
23	GCATTACAAT	ATGCTAAAAA	TTTTCAGCCA	TTTGCCCTAA	ATCATCAAAA	AGACATTCAG	300
	GTTTTGATGG	GAAGCCTTGT	GTACCTGAGA	CAAGGGATTG	AGAACTCACC	ATATGTTCAC	360
30	CTACTTGATG	CAAACCAGTG	GGCTGATATC	TGTGACATCT	TTACACGGGA	TGCTTGTGCC	420
	CTCCTGGGGC	TCTCCGTGGA	GTCCCCTCTC	AGTGTCAGTT	TCTCAGCAGG	TTGTGTGGCG	480
35	CTGCCAGCTT	TAATTAACAT	CAAAGCCGTG	ATTGAACAGA	GGCAGTGTAC	TGGAGTTTGG	540
33	AACCAGAAAG	ATGAATTACC	TATTGAAGTG	GACCTTGGTA	AAAAGTGCTG	GTATCACTCT	600
	ATATTTGCCT	GCCCCATTCT	TCGTCAGCAA	ACAACAGATA	ACAATCCACC	CATGAAATTG	660
40	GTCTGTGGTC	ATATTATATC	AAGAGATGCC	CTGAATAAAA	TGTTTAATGG	TAGCAAATTA	720
	AAATGTCCCT	ACTGTCCAAT	GGAACAAAGT	CCAGGAGATG	CCAAACAGAT	ATTTTTCTGA	780
45	AGAGATAACT	TTAGTTTGCA	ATTTGTAAGT	GAAACTGAAT	CGTGGGTGCA	TTTCAGAAGA	840
73	GAACGTTCCA	TATAATGCAG	CTAACCAAGG	ACTCCTGTGT	TTCTATAAGC	TAATGCTCCA	900
	GAAACTTTGC	CAACCTGTTA	GTGTACACAC	ACTGAGGGGA	GTGCTCCCGG	TGAATATTAT	960
50	CATAGGGCTT	TATTATATTC	TTGGTCTTCA	TTTCTGATCA	AGTAAATACA	CCAGCAGTTG	1020
	TCATTCAATG	CAGGTTTTTG	TACTTAATTA	TATGGTGATT	TTTTTACTTT	TTAAGAGCAG	1080
55	AAACGGAAAT	TGACCTCCCC	GCCATGTGTT	TAATATTCCT	CCTGCTTTTA	CTTTTGTCAT	1140
JJ	TTTCTTGATA	ATCGTAAGCC	TTGAGAGTGT	TTGTGAAAAA	GTTTTATTTC	CTGTTATGTA	1200
	ТАСАТААТТА	AATGAAAATT	CTTCAGAAAA	AGTTTGATAA	ATTGAATTGT	GGTTATGAAA	1260
60	CTAATTTGCA	TTTTTATTIG	CTTAAGAAAG	AAAGCTGTGA	TAGATTCCAG	ATATGCTTTT	1320

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	TGATGTTTTC	CTCTGCTCCA	GCTCCAAGAA	GTCAGCACAC	CTGCATTTTA	GCTCTGCATG	1380
_	CAGCCCCAGC	AGGCTGCGTG	TTTAAGAATT	TCATTGTTTA	ACTGGCTGGT	GTGAGAAGTC	1440
5	TTCCGTTAGC	ATAGAGTGGA	AGGAGTACTA	TIGTTIGGIT	GGGTTTTTGT	TTGTTTGTTT	1500
	TTTGTTTTTG	CTTTTATTGC	CAAGAGGTGC	TTGTTTTAAA	AGTATGTTTA	ATAAAATGAA	1560
10	ATTCTAAAGT	TAARAAGTGT	TCTTAAAGTT	GATATTTAAC	TCT		1603

15 (2) INFORMATION FOR SEQ ID NO: 38:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 1089 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: double
- (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 38:

25	GGCACGAGCT ACCTTTCTGC CTGCTTTGCT GGCTGCAACA GCACGAATCT CACGGGCTGT	60
	GCGTGCCTCA CCACCGTCCC TGCTGAGAAC GCAACCGTGG TTCCTGGAAA ATGCCCCAGT	120
20	CCTGGGTGCC AAGAGGCCTT CCTCACTTTC CTCTGTGTGA TGTGTATCTG CAGCCTGATC	180
30	GGTGCCATGG CAAGACACCC TCAGTCATCA TCCTCATCAG GACAGTCAGC CCTGAACTCA	240
	AGTCTTACGC TTTGGGAGTT CTTTTTCTCC TCCTTCGTTT GTTGGGCTTC ATCCCTCCAC	300
35	CCCTCATCTT CGGGGCTGGC ATCGACTCCA CCTGCCTGTT CTGGAGCACG TTCTGTGGGG	360
	AGCAAGGCGC CTGCGTCCTC TACGACAATG TGGTCTACCG ATACCTGTAT GTCAGCATCG	420
40	CCATCGCGCT CAAATCCTTC GCCTTCATCC TGTACACCAC CACGTGGCAG TGCTGAGGAA	480
40	AAACTATAAA CGCTACATCA AAAACCACGA GGGCGGGCTG AGCACCAGTG AGTTCTTTGC	540
	CTCTACTCTG ACCCTAGACA ACCTGGGGAG GGACCCTGTG CCCGCAAACC AGACACATAG	600
45	GACAAAGTTT ATCTATAACC TGGAAGACCA TGAGTGGTGT GAAAACATGG AGTCCGTTTT	660
	ATAGTGACTA AAGGAGGGCT GAACTCTGTA TTAGTAATCC AAGGGTCATT TTTTTCTTAA	720
	AAAAAGAAAA AAAGGTTCCA AAAAAAACCA AAACTCAGTA CACACACAC GGCACAGATG	780
50	CACACACAC CAGACAGACA CACCGACTTT GTCCTTTTTC TCAGCATCAG AGCCAGACAG	840
	GATTCAGAAT AAGGAGAGAA TGACATCGTG CGGCAGGGTC CTGGAGGCCA CTCGCGCGGC	900
55	TGGGCCACAG AGTCTACTTT GAAGGCACCT CATGGTTTTC AGGATGCTGA CAGCTGCAAG	960
	CAACAGGCAC TGCCAAATTC AGGGAACAGT GGTGGCCAGC TTGGAGGATG GACATTTCTG	1020
60	GATACACATA CACATACAAA ACAGAAAACA TTTTTTAAAA GAAGTTTCCT AAAATAAAAA	1080

	AAAAAAA	1089
5	(2) INFORMATION FOR SEQ ID NO: 39:	
10	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 629 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 39:	
	AGCTCAGTTC CCTTAGAAAT GAAATTTTAA ATGACACTAC CAGGTAAGCC ACTGAGACCA	60
	GTGGAGGTGA TAGCTAAGAA CATAAGGAAT TAAGAATTTT TAATGGAGAA AGGAGGTAAT	120
20	GAATACCAGT TACATCCTAA GACTCACTGT AGTGGTGAGT GTTGTAATTT ATCTCGCTAT	180
	CCATCCTCTT TTAAGTTTTT CCTTAGAAAG TCCTCTATTG GTACCTTGGA GGGACTGCTG	240
25	TCAAAATATA TGGAAAAGTG GGTCTGTGTG GTACAAGAGG TGGACTTTGC CACACATGGA	300
23	AGTTTGCTGC CAAGATCTTC ACTAATGAAA GAAATCACCA GTGAGCTGCA CAGATTAGCC	360
	AAATACTGAG CTCATTAGAA CTACTAAGGC CTGGACATTT CTGCCTAATC CAGGACTCCT	420
30	GTAATTATCA GTCTTTGCTT TGGAGCTTCC CATTGTGTAG CTGARAATTT GTCATATCTG	480
	CATTATAATC TAAGGCTCCA CATACTTAAT CCTGCTTCTC CCCCTTTTTC TTTCCCTTTC	540
25	CCAGCGGTCA GCTCTGCTGC ATAGTCTGAA GACTTTCCCT GCCCAATCCT GATAAAATTC	600
35	TTGCACTCGT AACCCCATCT CAGTGTCTG	629
40	(2) INFORMATION FOR SEQ ID NO: 40:	
45	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1964 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 40:	
50	AAGAAGACAT GGAAATTGCT GAAGGATGTT TCAGGCATAT TAAGAAAATC TTTACGCAGC	60
	TTGAGGAATT CAGAGCCTCT GAATTGCTTC GAAGTGGACT GGACAGATCT AAATACCTTT	120
55	TAGTGAAAGA AGCCAAAATT ATTGCTATGA CCTGTACTCA TGCTGCCTTA AAACGACATG	180
	ACTTGGTCAA GCTAGGTTTC AAGTATGACA ACATTTTGAT GGAAGAGGCT GCTCAGATTC	240
60	TGGAGATAGA AACTTTTATC CCTCTTCTTC TACAGAATCC TCAGGATGGA TTTAGCCGAC	300

	TAAAACGATG GATTATGATT GGCGATCATC ACCAGTTACC TCCAGTTATT AANGAACATG	360
	GCCTTTCAAA AGTACTCAAA CATGGAGCAG TCTCTCTTCA CTCGCTTTGT TCGCGTTGGA	420
5	GTTCCGACTG TTGACCTTGA TGCTCAAGGG AGAGCCAGAG CAAGCTTGTG CAMCTNCTAC	480
	AACTGGCGAT ACAAGAATCT AGGAAACTTA CCCCATGTGC AGCTCTTGCC AGAGTTTAGT	540
	ACAGCAAATG CTGGCTTACT GTATGACTTC CAGCTCATTA ATGTTGAAGA TTTTCAAGGA	600
10	GTGGGAGAAT CTGAACCTAA TCCTTACTTC TATCAGAATC TTGGAGAGGC AGAATATGTA	660
	GTAGCACTTT TTATGTACAT GTGTTTACTT GGTTACCCTG CTGACAAAAT CAGTATTCTA	720
15	ACAACATATA ATGGCCAAAA GCATCTTATT CGCGACATCA TCAATAGACG ATGTGGAAAC	780
	AATCCATTGA TTGGAAGACC AAACAAGGTG ACAACTGTTG ATAGATTTCA AGGTCAACAG	840
20	AATGACTATA TTCTTCTTTC TCTGGTACGA ACCAGGGCAG TGGGCCATCT GAGGGATGTC	900
20	CGTCGCTTGG TAGTGGCCAT GTCTAGAGCC AGACTTGGAC TTTATATCTT CGCCAGAGTA	960
	TCCCTCTTCC AAAACTGTTT TGAACTGACT CCAGCTTTCA GTCAGCTCAC AGCTCGCCCC	1020
25	CTTCATTTGC ATATAATTCC AACAGAACCT TTCCCAACTA CTAGAAAGAA TGGAGAGAGA	1080
	CCATCTCATG AAGTACAAAT AATAAAAAAT ATGCCCCAGA TGGCAAACTT TGTATACAAC	1140
20	ATGTACATGC ATTTGATACA GACTACACAT CATTATCATC AGACTTTATT ACAACTACCA	1200
30	CCTGCTATGG TAGAAGAGGG TGAGGAAGTT CAAAATCAAG AAACAGAATT GGAAACAGAA	1260
	GAAGAGCCCA TGACTGTTCA AGCTGACATC ATACCCAGTC CAACAGACAC CAGCTGCCGT	1320
35	CAAGAAACTC CAGCCTTTCA AACTGACACC ACCCCCAGTG AGACAGGAGC CACTTCCACT	1380
	CCAGAAGCCA TCCCTGCTTT ATCTGAGACC ACCCCTACTG TGGTAGGAGC TGTATCTGCA	1440
40	CCGGCAGAAG CTAACACACC TCAGGATGCC ACATCTGCCC CAGAAGAGAC CAAGTAGCCA	1500
40	AACTGTAGTC CTTCTAAAGG AGGACATGGC AGTCAAAAAG TCTGAGTAAA GCTGTTTTTT	1560
	GTATTTATA TITGCTTCTG CCATTTTACT GTCACTAATT AATGTTTAGT TCTTATATTT	1620
45	GTTAACTGAT TICGGTGTCT TGAATATATT TTTTTAAATT ATGTGTATGA ACAATTCTAG	1680
	TTTCATTTGT TCAATCAGAA GAGCAAATAA CCATTCCTTT CATGTTTTGA TCACTGAGTG	1740
50	TGTCTGTAAT CATACCTACA TTAAAATCAT TTTCTATGAA TATATAATAT ATACTTCACA	180
50	TTTTTAGTGA ACTTCTCTAA AGAAGAGGAC AGAATATACT GGACTTAACC ACGAATACCC	186
	TTGAGTGTCC AAATTGGGAA GGAACTKGTT TCTTCYGTTA TACTAYCAAA TGCTTAAATT	192
55	CKGTTTCCTT TTTTCTTACC TTTGTTTGCT GTCTTTATGT AAAG	196

60 (2) INFORMATION FOR SEQ ID NO: 41:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1522 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 41:

10	CGTGTCCGCG	CGCCTGGGAG	ACGCTGCCTC	GGCCCGGACG	CGCCCGCGCC	CCCGCGGCTG	60
	GAGGGTGGTC	GCCACTGGGA	CACTGTGAAC	CAGGAGTRAG	TCGGAGCTGC	CGCGCTGCCC	120
15	AGGCCATGGA	CTGTGAGGTC	AACAACGGTT	CCAGCCTCAG	GGATGAGTGC	ATCACAAACC	180
13	TACTGGTGTT	TGGCTTCCTC	CAAAGCTGTT	CTGACAACAG	CTTCCGCAGA	GAGCTGGACG	240
	CACTGGGCCA	CGAGCTGCCA	GTGCTGGCTC	CCCAGTGGGA	GGGCTACGAT	GAGCTGCAGA	300
20	CTGATGGCAA	CCGCAGCAGC	CACTCCCGCT	TGGGAAGAAT	AGAGGCAGAT	TCTGAAAGTC	360
	AAGAAGACAT	CATCCGGAAT	ATTGCCAGGC	ACCTCGCCCA	GGTCGGGGAC	AGCATGGACC	420
25	GTAGCATCCC	TCCGGGCCTG	GTGAACGGCC	TGGCCCTGCA	GCTCAGGAAC	ACCAGCCGGT	480
	CGGAGGAGGA	CCGGAACAGG	GACCTGGCCA	CTGCCCTGGA	GCAGCTGCTG	CAGGCCTACC	540
	CTAGAGACAT	GGAGAAGGAG	AAGACCATGC	TGGTGCTGGC	CCTGCTGCTG	GCCAAGAAGG	600
30	TGGCCAGTCA	CACGCCGTCC	TTGCTCCGTG	ATGTCTTTCA	CACAACAGTG	AATTTTATTA	660
	ACCAGAACCT	ACGCACCTAC	GTGAGGAGCT	TAGCCAGAAA	TGGGATGGAC	TGAACGGACA	720
35	GTTCCAGAAG	TGTGACTGGC	TAAAGCTCGA	TGTGGTCACA	GCTGTATAGC	TGCTTCCAGT	780
	GTAGACGGAG	CCCTGGCATG	TCAACAGCGT	TCCTAGAGAA	GACAGGCTGG	AAGATAGCTG	840
	TGACTTCTAT	TTTAAAGACA	ATGTTAAACT	TATAACCCAC	TTTAAAATAT	СТАСАТТААТ	900
40	ATACTTGAAT	GAAAATGTCC	ATTTACACGT	ATTTGAATGG	CCTTCATATC	ATCCACACAT	960
	GAATCTGCAC	ATCTGTAAAT	CTACACACGG	TGCCTTTATT	TCCACTGTGC	AGGTTCCCAC	1020
45	TTAAAATTA	AATTGGAAAG	CAGGTTTCAA	GGAAGTAGAA	ACAAAATACA	ATTTTTTTGG	1080
	ТАААААААА	TTACTGTTTA	TTAAAGTACA	ACCATAGAGG	ATGGTCTTAC	AGCAGGCAGT	1140
	ATCCTGTTTG	AGGAAAGCAA	GAATCAGAGA	AGGAACATAC	CCCTTACAAA	TGAAAAATTC	1200
50	CACTCAAAAT	AGGGACTATC	YATCTTAATA	CTAAGGAACC	AACAATCTTC	CTGTTTAAAA	1260
	AACCACATGG	CACAGAGATT	CNGAACTAAA	GTGCTGCACT	CAAATGATGG	GAAGTCCCGG	1320
55	CCCCAGTACA	CCAGGGGCTT	TGGACTTTTT	TCAACTTCGT	TTCCTTTTGT	TTGGANTCCA	1380
	AAAGAACCAC	TTTGTGGTTC	TTAAAAGGGT	GTGAAGGTGA	TTTAAGGGGC	CCAGGTCAGC	1440
	CACTGGTTGG	TTTACAAAAT	CNGGGTAACT	AACTGCATAC	AACTITTTCC	CNTTTCCATG	1500
60	NCATCAGGAC	TTTGCTAAAG	AC				1522

5	(2) INFORMATION FOR SEQ ID NO: 42:	
10	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 875 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 42:	
15	TGGGATTTCC CTTTATCATG GAGGCCTTGT CCCACTTCCT CTATGTCCCT TTCCTTGGTG	60
	TCTGTGTCTG TGGGGCCATC TACACTGGCC TGTTCCTTCC TGAGACCAAA GGCAAGACCT	120
20	TCCAAGAGAT CTCCGAGGAA TTACACAGAC TCAACTTCCC CAGGCGGGCC CAGGGCCCCA	180
20	CGTGGAGGAG CCTGGAGGTT ATCCAGTCAA CAGAACTCTA GTCCCAAAGG GGTGGCCGTA	240
	GCCAAAGCCA GCTACCGTCC TGTCCTCTGC TTCCTGCCAG GGCCCTGGTC CTCAMTYCCT	300
25	YCTGCATTCC TCATTTAAGG AGTGTTTATT GAGCACCCTT TGTGTGCAGA CATGGCTCCA	360
	GGTGCTTAGC AATCAWTGGT GAGCGTGGTA TCCAGGCTAA AGGTAATTAA CTGACAGRAA	420
20	ATCAGTAACA ACATAATTAC AGGYTGGTTG TGGCAGYTCA TGACTGTAAT CCCAGCACTT	480
30	TTGGGAGCCA AGGTGGGARG ATCAATTGAG GCCAGAGTTT GAAAMCAGCT AGGTAACATA	540
	GTGAGACCCC CTATCTCTAC AAAAAATTTT AAACATTAGC TGGGCATGGT GGTATGTGCT	600
35	AACAGCTCTA GCTACTCAGG AGGCTGAGGC AGCAGGATCA CTTGAGTCCA AGAGTTCAAG	660
	GTAGCAGTAA GCTACAATCA CACCACTGCA TGCCAGACTG GGTGACAGAG GGAGACTTCA	720
40	TCTCTTTAAA ACATAATAAT AATAATTACA GACTCAGGAA ATGCAGTGAA AGAAAAATAC	780
40	AGGTTGGCCA GGTGAGGTGG CTGATGCCTG TAATCCCAGC ACTTTGGGAG GCCAAGATGG	840
	GAAGATTGCT TTGAGACCAG AAGTTTGAGA CCAGC	875
45		
	(2) INFORMATION FOR SEQ ID NO: 43:	
50	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 843 base pairs(B) TYPE: nucleic acid	
55	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
55	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 43:	
	CCCACGCGGT CCGNATCGTC CTTCCCTCAC TTCAGAGGGT GGCCAGAGCT GAATACCCAG	60
60	AGAGGGACAA GTAAGGGTCC AGTTCCAAAA CATCATGAGG ATGTATCATC CCACGTGTCT	120

	CACCTGACAG TTACAGAGGA AACCCGCACC CAGAATGCAC GTGCTGTCTT ATGGGAACAC	180
5	TCAGCGCAGA GTGCTCAGGT CCGGCCACAC TCGGGCTGTG CTTGGTCGTG CCATGGAATT	240
	CCTCAGGACT TTCTCAGCCT CCCTAATGGC AGAAGCCCCT TTACAGCAAG ACATTTACCG	300
	TTTGTCTGAA AATAGCCGAA CTGAGCTTTT CTTCAGGCTA TATGAGAAGT CTCTAGACAG	360
10	TGGGCACCGT CAGAAAGCCC AGAGCCTTGT GATAGCTCCC ACCCTGCCTG GCTCAGATCT	420
	TCCCATTTT TTTCCTCTGG CACTAACCTC ACCTTTTGTT TTTTTGTGTT TGTGTTTGTT	480
15	TTTGTTTTTG CAGAGTTGGA TTACAGAAAC TCCTATGAAA TTGAATATAT GGAGAAAATT	540
	GGCTCCTCCT TACCTGTAAG TTCGTCTGCC TCGGGCCACT TAGGGGACTC GCTTTCCTGC	600
	CTTCAGGGGC CTCCTCCCCT GTGCAGAGTG TCTCTGGGAG CTCAGACCCC AAATCGAGTG	660
20	TTTTCTGTGT ACACAGCTTC CCGGGTGCAC AGCAATGATG GACTGGGGCT GGGGGGTTGA	720
	GGTTTGTACT CAATCCACTT CGTTTGACAT TTTCAGGGAG AAAATGATAG AATACAATTA	780
25	GACGTCCTGC AGAATTACTT TCCTAGACTG AGAAAGAGCT AGAGATTTCT TTAAAAAAAA	840
	AAA	843
30	(2) INFORMATION FOR SEQ ID NO: 44:	
35	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 489 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 44:	
	CTCTTAGGCT TTGAAGCATT TTTGTCTGTG CTCCCTGATC TTCAGGTCAC CACCATGAAG	60
	TTCTTAGCAG TCCTGGTACT CTTGGGAGTT TCCATCTTTC TGGTCTCTGC CCAGAATCCG	120
45	ACAACAGCTG CTCCAGCTGA CACGTATCCA GCTACTGGTC CTGCTGATGA TGAAGCCCCT	180
	GATGCTGAAA CCACTGCTGC TGCAACCACT GCGACCACTG CTGCTCCTAC CACTGCAACC	240
50	ACCGCTGCTT CTACCACTGC TCGTAAAGAC ATTCCAGTTT TACCCAAATG GGTTGGGGAT	300
	CTCCCGAATG GTAGAGTGTG TCCCTGAGAT GGAATCAGCT TGAGTCTTCT GCAATTGGTC	360
	CTCCCGAATG GTAGAGTGTG TCCCTGAGAT GGAATCAGCT TGAGTCTTCT GCAATTGGTC ACAACTATTC ATGCTTCCTG TGATTTCATC CAACTACTTA CCTTGCCTAC GATATCCCCT	360 420

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	(2) INFORMATION FOR SEQ ID NO: 45:	
5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 534 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 45:	
	GAAGCAGTGT GTATCTATGA TTATATCTCT GTTCATCTAT ATATTTTTGA CATGTAGCAA	60
15	CACCTCTCCA TCTTATCAAG GAACTCAACT CGGTCTGGGT CTCCCCAGTG CCCAGTGGTG	120
15	GCCTTTGACA GGTAGGAGGA TGCAGTGCTG CAGGCTATTT TGTTTTTTGT TACAAAACTG	180
	TCTTTTCCCT TTTCCCCTCC ACCTGATTCA GCATGATCCC TGTGAGCTGG TTCTCACAAT	240
20	CTCCTGGGAC TGGGCTGAGG CAGGGGCTTC GCTCTATTCT CCCTAACCAT ACTGTCTTCC	300
	TTTCCCCTTG CCACTTAGCA GTTATCCCCC CAGCTATGCC TTCTCCCTCC CTCCCTTGCC	360
25	CTGGCATATA TTGTGCCTTA TTTATGCTGC AAATATAACA TTAAACTATC AAGTGAAAAA	420
23	AAAAAAAAA AAAACTCCAA GGGGGGCCG GTACCCAATT CCCCCTATAN TGAGTCNTAT	480
	TACAATTCAC TGGGCCGTCG TTTTACAACG TCGTGAATGG GAAAACCTGG GCGT	534
30		
	(2) INFORMATION FOR SEQ ID NO: 46:	
35	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1374 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 46:	
	GGCACGAGTC CGGGATGAGC TCAGCCGCGG CCGACCACTG GGCGTGGTTG CTGGTGCTCA	60
45	GCTTCGTGTT TGGATGCAAT GTTCTTAGGA TCCTCCTCCC GTCCTTCTCA TCCTTCATGT	120
	CCAGGGTGCT GCAGAAGGAC GCGGAGCAGG AGTCACAGAT GAGAGCGGAG ATCCAGGACA	180
50	TGAAGCAGGA GCTCTCCACA GTCAACATGA TGGACGAGTT TGCCAGATAT GCCAGGCTGG	240
50	AAAGAAAGAT CAACAAGATG ACGGATAAGC TCAAAACCCA TGTGAAAGCT CGGACAGCTC	300
	AATTAGCCAA GATAAAATGG GTGATAAGTG TCGCTTTCTA CGTATTGCAG GCTGCCCTGA	360
55	TGATCTCACT CATTTGGAAG TATTATTCTG TCCCTGTGGC TGTCGTGCCG AGTAAATGGA	420
	TAACCCCTCT AGACCGCCTG GTAGCCTTTC CTACTAGAGT AGCAGGTGGT GTTGGAATTA	480
60	CCTGTTGGAT TTTAGTCTGT AACAAAGTTG TCGCTATTGT GCTTCATCCG TTCAGCTGAA	540

	CAGGAGGATG	GATACAGCCG	CGAGGCTAAA	AAACGGATTT	CCTCTTCCTA	GCTTAAAATC	600
	TGATTTACAC	TGTTTTGTTT	TTTAAGAAAC	AAAAGTGCAT	AGTTTAGATT	TTTTTTTTTG	660
5	TTGAATATGT	TTGTTCTTGG	ACTITATGAG	AGAGTCTTAT	AAGAATCACG	ATTTTCTACA	720
	CCTGTCATTG	AGCCAAGAAA	GTCCAGTTTA	TGACACGTAT	GTACTAGTGA	ACACCGTCCT	780
10	CGATCTGTAC	GAAATGTGAA	ATGTTTAGGG	ACATCTCCAT	GCTGTCACTT	GTGATTTGCC	840
	CTCTTATGTA	TTTTGGTCAT	ATTGCCAACT	GGAAAGTCAA	AATTTTCTAA	CAACTTTAAG	900
	TAAGTTCTTT	GAAGACTTAG	TGCTGTTTTT	AATCCAGTTT	AGAAAGTAAC	ТТААТТТТАА	960
15	TACCACTACT	AAAAATTCGA	AAATTTCTTC	TTTAATCACA	TTCAATATGG	TTAAAAGAAC	1020
	AACACTAATT	GACATTGCGT	GGGCTTTTTC	TCCCTTTGTT	TAAAATGTCA	TTTGTTGAGC	1080
20	AAGAGTTGTA	TAGTATTATC	TACTTACTTG	AGGCTGTTAA	TTTTTCATTA	CAGTGTTTTG	1140
	TAAATGTATC	CACGAGACCA	TGATGCATTG	TTTTGTGCTC	AACTTGTGTT	TTGTATTTAA	1200
	AGCATTTTGA	ATGAAGTGTA	TTTTATAAGC	ATTTAATATT	TATGCTCTTT	AGAATGGAAC	1260
25	ACAGAAAACA	AACCTTATAA	GTCCTGATTA	ATCTGAACCA	ATAACCTGTG	TGGCCTACAA	1320
	AGTATAATTC	TATTAAATGT	TCCTTAAAAC	AAAAAAAA	AAAAAAAA	AAAA	1374
30							
	(2) INFORMA	ATION FOR SE	O ID NO: 47	' <u>.</u>			
35		(B) TYPI (C) STRA	_	ICS: se pairs acid double			
35 40	(i)	SEQUENCE CH (A) LENG (B) TYPI (C) STR	HARACTERIST: GTH: 596 base E: nucleic and ANDEDNESS: O DLOGY: linea	ICS: se pairs acid double ar	: 4 7:		
	(i)	SEQUENCE CH (A) LENC (B) TYPI (C) STRA (D) TOPO SEQUENCE I	HARACTERISTI GTH: 596 base E: nucleic and ANDEDNESS: O DLOGY: lineand DESCRIPTION:	CCS: se pairs acid double ar : SEQ ID NO:		TTCATTTACT	60
40	(i)	SEQUENCE CH (A) LENC (B) TYPH (C) STRA (D) TOPC SEQUENCE I	HARACTERIST: GTH: 596 ba: E: nucleic a ANDEDNESS: 6 DLOGY: linea DESCRIPTION: TGGACATGAA	ICS: se pairs acid double ar : SEQ ID NO:	TTCAAGTTTA		60 120
	(i) (xi) GAATTCGNCA AAGTTAGTTA	SEQUENCE CH (A) LENC (B) TYPH (C) STRA (D) TOPC SEQUENCE I CGAGATTACT AATCATGTGC	HARACTERIST: GTH: 596 ba: E: nucleic a ANDEDNESS: 6 DLOGY: linea DESCRIPTION: TGGACATGAA CTTCCATGAG	ICS: se pairs acid double ar : SEQ ID NO: AGAACTCAGG	TTCAAGTTTA GTAACTTGGA		120
40	(i) (xi) GAATTCGNCA AAGTTAGTTA	SEQUENCE CH (A) LENC (B) TYPH (C) STRA (D) TOPC SEQUENCE I CGAGATTACT AATCATGTGC GTCATATATA	HARACTERIST: STH: 596 ba: E: nucleic a ANDEDNESS: 0 DLOGY: line DESCRIPTION: TGGACATGAA CTTCCATGAG TTCTACACTG	ICS: se pairs acid double ar : SEQ ID NO: AGAACTCAGG CCTTCATTTG	TTCAAGTITA GTAACTTGGA GACCAAAGGG	AAATGGAAAT ATTATAGATT	120
40	(xi) (xi) GAATTCGNCA AAGTTAGTTA AATAACACTA	SEQUENCE CH (A) LENC (B) TYPH (C) STRA (D) TOPC SEQUENCE I CGAGATTACT AATCATGTGC GTCATATATA TCATTCCTGC	HARACTERIST: STH: 596 ba: E: nucleic and state of the sta	ICS: se pairs acid double ar : SEQ ID NO: AGAACTCAGG CCTTCATTTG CTACCATATG	TTCAAGTITA GTAACTTGGA GACCAAAGGG ATTTCATTGA	AAATGGAAAT ATTATAGATT AGAAAAGTCC	120 180
40 45	(xi) GAATTCGNCA AAGTTAGTTA AATAACACTA ACAATCACCA TTACATTTAT	SEQUENCE CHECK (A) LENG (B) TYPH (C) STRAIN (D) TOPK (D) TOPK (D) TOPK (D) SEQUENCE IN CGAGATTACT (CATATATA (CATTCCTGC (CCTTTTCCTA (C) (CCTTTTCCTA (C)	HARACTERIST: STH: 596 ba: E: nucleic and properties of the colory of the colors of the colory of the colors of the colory of the colory of the colors of the	ICS: se pairs acid double ar : SEQ ID NO: AGAACTCAGG CCTTCATTTG CTACCATATG ATAGAAAACA GGGTAAACTA	TTCAAGTITA GTAACTTGGA GACCAAAGGG ATTTCATTGA ATAAATATAG	AAATGGAAAT ATTATAGATT AGAAAAGTCC	120 180 240
40 45	(xi) GAATTCGNCA AAGTTAGTTA AATAACACTA ACAATCACCA TTACATTTAT ACCCTTATTA	SEQUENCE CHECK (A) LENG (B) TYPH (C) STRAME (D) TOPK (D) TOPK (D) TOPK (D) SEQUENCE IN CGAGATTACT (CATATATA (CATTCCTGC (CCTTTTCCTA (CTTATTATA)	HARACTERIST: STH: 596 ba: E: nucleic andedness: oblogy: line; DESCRIPTION: TGGACATGAA CTTCCATGAG TTCTACACTG TGACAGGTAT ATATCTGCAT TTCAATGTGA	ICS: se pairs acid double ar : SEQ ID NO: AGAACTCAGG CCTTCATTTG CTACCATATG ATAGAAAACA GGGTAAACTA GAACTGCTGC	TTCAAGTITA GTAACTTGGA GACCAAAGGG ATTTCATTGA ATAAATATAG AGAAAAAATA	AAATGGAAAT ATTATAGATT AGAAAAGTCC TCATTAGAAA	120 180 240 300
40 45	(xi) GAATTCGNCA AAGTTAGTTA AATAACACTA ACAATCACCA TTACATTTAT ACCCTTATTA	SEQUENCE CHECK (A) LENG (B) TYPH (C) STRAIN (D) TOPK (D)	HARACTERIST: STH: 596 ba: E: nucleic andedness: oblogy: line; DESCRIPTION: TGGACATGAA CTTCCATGAG TTCTACACTG TGACAGGTAT ATATCTGCAT TTCAATGTGA ATATTCATAA	ICS: se pairs acid double ar : SEQ ID NO: AGAACTCAGG CCTTCATTTG CTACCATATG ATAGAAAACA GGGTAAACTA GAACTGCTGC ATTTCAAAT	TTCAAGTITA GTAACTTGGA GACCAAAGGG ATTTCATTGA ATAAATATAG AGAAAAAATA CATTGAAAAT	AAATGGAAAT ATTATAGATT AGAAAAGTCC TCATTAGAAA TGCTTTATAA TACCTTAAAA	120 180 240 300 360
40 45	(xi) GAATTCGNCA AAGTTAGTTA AATAACACTA ACAATCACCA TTACATTTAT ACCCTTATTA TATTTTCTTG	SEQUENCE CHECK (A) LENG (B) TYPH (C) STRAIN (D) TOPK (D)	HARACTERIST: STH: 596 ba: E: nucleic andedness: oblogy: line; DESCRIPTION: TGGACATGAA CTTCCATGAG TTCTACACTG TGACAGGTAT ATATCTGCAT TTCAATGTGA ATATTCATAA TACTCATATA	ICS: se pairs acid double ar : SEQ ID NO: AGAACTCAGG CCTTCATTTG CTACCATATG ATAGAAAACA GGGTAAACTA GAACTGCTGC ATTTCAAAT ACAGTATAAA	TTCAAGTITA GTAACTTGGA GACCAAAGGG ATTTCATTGA ATAAATATAG AGAAAAAATA CATTGAAAAT ATTCCTATGT	AAATGGAAAT ATTATAGATT AGAAAAGTCC TCATTAGAAA TGCTTTATAA TACCTTAAAA CAATCTCTTT	120 180 240 300 360 420

5	(2) INFORMATION FOR SEQ ID NO: 48:	
10	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 851 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 48:	
15	CACATGAAGA CACACAGTGG TGAGAAGCCC TTCCGCTGCG CCCGCTGTCC TTATGCCTCT	60
	CCTCATCTGG ATAACCTGAA ACGGCACCAG CGCGTCCATA CAGGAGAGAA GCCCTACAAG	120
20	TGCCCCCTCT GCCCTTATGC CTGTGGCAAT CTGGCCAACC TCAAGCGTCA TGGTCGCATC	180
20	CACTCTGGTG ACAAACCTTT TCGGTGTAGC CTTTGCAACT ACAGCTGCAA CCAGAGCATG	240
	AACCTCAAAC GTCACATGCT GCGGCACACA GGCGAGAAGC CTTCCGCTGT GCCACCTGCG	300
25	CCTATACCAC GGGCCACTGG GACAACTACA AGCGCCACCA GAAGGTGCAT GGCCACGGTG	360
	GGGCAGGAGG GCCTGGTCTC TCTGCCTCTG AGGGCTGGGC CCCACCTCAT AGCCCACCCT	420
20	CTGTTTTGAG CTCTCGGGGC CCACCAGCCC TGGGGACTGC TGGCAGCCGG GCTGTCCACA	480
30	CAGACTCATC CTGAACTAGG TCCTTCTTCC CCATGTTTTA TACAGACGGA CCAGAAGCCA	540
	CCTTTTTCTC CCCCGCTGGC CAGGGGCTCC ACACAGACTA ACGTAGGCAC TATAAGGACC	600
35	AGCCCAACCC CATGGGCGGG GGGGCCCATA TGGACCAGGG GACCTTGCCT TGACTGAGGC	660
	ACTTCACGAG CTCAGTGAGA AGGGCCCTGT ATTCACCTCC ACTGCCCCCA GGGGCTGTGG	720
40	ACAAACCGGC TGGGGGACTG CCCAGCCTCC CACCTGTTTA TTTAACTTAT TTCAGTGCTT	780
40	TATAATAAAG GAAACACTAA CAAAGCCATG TCTATGCTGA ATTGGCAATG GCAGGCAATT	840
	TGGCCTTACC C	851
45		
5 0	(2) INFORMATION FOR SEQ ID NO: 49:	
50	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 2020 base pairs	
	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
55	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 49:	۰
	GTGAAATGAA AACAGTCTTT TTATAGCCTT TAGCTTGTGA GTTTGGAAGT TTGGGGGGTC	12
40		12

	ACACAGACCC	AGGTGAACAC	GCTGACTGTG	AACCTGCCCT	GTATCCGGAG	CTGTGCTGGG	180
5	CACTGAGGGG	ATGCAACAAA	ATTAGGAGAG	GWTCCTTGCT	CCCAACGTCT	ACTTCTCCTA	240
	CCTCAACAGG	GGTCCAGGGT	GCAGTGAACT	CAGTTCTTGG	CCCTTGGGTG	AGGATTCATG	300
	GATGAATGAA	AGCTAGACCT	GATGGGGAGG	CATTATGACT	AAATAGGCCC	AGCCTCCTTC	360
10	CCTTCCAGCT	CTGTCCTAGG	AGCATAGGCG	GGAAATCTGA	GTAGAGTCTG	ACTGCAGTTT	420
	TTGCTTATGA	TTTGTAAAAG	CCGTCATGGG	GTCAATAAGA	AAATAGGGGT	GATGGAGGG	480
15	GAGAAGCCCA	GGACTGGGAG	AATCGCACGT	GCCCCAGGGG	TTTTCACCAA	GGATTTTCAA	540
13	GACAAACTGG	AGTAAGAATT	AAAGCCCCAG	AGGATTTAAT	TATCCTGGTT	TGCAAAAGAG	600
	CCTCCCATGC	CAGTACCGCC	CAGCCTTGGA	GGCCGGAATG	CTCATGGCCC	CTGTGGTCTG	660
20	CTTGTCCTTC	AGCCCATGCC	CAGCAGATAC	CTCTCTGACT	GGAGACGGGC	TCAAAGCTGG	720
	ATTAGAAAGG	GGAGMGGCAC	TTGTGACTTT	GTTTGACTCT	GTGACTCACT	TCCTCGCTCA	780
25	CACCTTGTTT	GAACTACTGG	ACTTTCAACT	GGCTTTCCTT	AGGTCAGGCA	AGCAGACAGC	840
	TCCCCACTGA	AGAGGTCTGT	ACAGTGACAA	ccceeccee	CAGCAAGGAC	ACAGATGCAG	900
	CCACAGTAAG	GCTCCATCAG	GACTGGGTCA	GTGATGGCAA	CAGGATGGCC	AAGGATGGCT	960
30	CTAGAACAYT	CTGTCCATGC	GTCACTCCCC	CCAGTTTTRT	TTTTAGCTTT	GGCTTCAGGG	1020
	AGTGACAGCC	ATCACAAATA	GCCACATTCT	GCTCTACTCT	CCAACATACC	AGATTSTACA	1080
35	CTGTTGTTAT	TTCATGAGAC	GTGAATGTTG	CAGAGAGTGG	GGGGATTCTG	GTTGTTAAGG	1140
	AACTTACACT	GGGGAGCTTT	ACTCTTCCGT	GTCAACAATG	TGACTACATG	TTCTCCAGAT	1200
	TAGCCACACA	TGCAAACATC	AGTGTCCTTC	TAGCTTTANC	CGAGAAAGAA	ACCAGTCCCA	1260
40	GGGAATGAAT	GGTGGTCTCC	CCACTCCCGG	CAGCACTTTA	GGCAGCCCAT	AAGCTATGCG	1320
	AGAATGTGAA	CGCTCACCTT	GCTCCGTCAC	GGTTCTGACC	TACCACATAA	ACAGGAAGAA	1380
45	GCCAGTGACC	GGAACAGCTC	TAGGAATAAC	AAGTCAGAAT	AGAAGTGTCC	TTTATATTAC	1440
	CAGAAAATAT	GGGCTTGGCC	TAAGTCGCTG	TCTCCTAACC	TGCCGGGGTC	ATTCCCCACC	1500
	AAACACCCCA	TACTAAGGAG	CCATGAGCCA	CCTGGACATT	CACCTTTTCT	TTGACCATCT	1560
50	GGAGTCTGGG	GCAACTTAAG	GAAGGCNCCA	CACAGTGGTG	CAGGCACATT	TCCAAGCGTA	1620
	GGTGTCCCTG	GCTTTTGTGG	CCAAAGCTAG	TGTTATGGTC	AACAACAGGC	CAGGGTCTGT	1680
55	GGGGCACTGA	CCTTGAAAGT	GGCAAAATGG	AGGTTTCACA	GGCTGTGCGG	GAGCAGGACG	1740
	GCTTGCTTCA	TCTAACAATC	TCAGTTTCCT	TTAAAAAAAG	AAAGAAAGGA	AAAGATTTCA	1800
	TAAGCAGGTG	TCAGTGGACA	GTTTAAGYAC	TTAACCATTT	CTCTTTCTTC	TTATGGATGT	1860
60	GAACTGTGCT	GTGGATAAAT	CATTIGTATT	TCTTGAATGT	TCTCTATGAC	TAACAGTTAT	1920

	TAAGTCGGTT GTGTATATGT GTAACTAATG TAACTGCCTT TTAAAATTTC ATTACAATAA	1980
5	AAATGACTTT GCTCTGAAMA AAAAAAAAA AAAAACTCGA	2020
10	(2) INFORMATION FOR SEQ ID NO: 50:	
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 2432 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double	
15	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 50:	
20	ATGAAGGGTC GTTGGTGGGA AAGATGGCGG CGACTCTGGG ACCCCTTGGG TCGTGGCAGC	60
20	AGTGGCGGCG ATGTTTGTCG GCTCGGGATG GGTCCAGGAT GTTACTCCTT CTTCTTTGT	120
	TGGGGTCTGG GCAGGGGCCA CAGCAAGTCG GGGCGGGTCA AACGTTCGAG TACTTGAAAC	180
25	GGGAGCACTC GCTGTCGAAG CCCTACCAGG GTGTGGGCAC AGGCAGTTCC TCACTGTGGA	240
	ATCTGATGGG CAATGCCATG GTGATGACCC AGTATATCCG CCTTACCCCA GATATGCAAA	300
20	GTAAACAGGG TGCCTTGTGG AACCGGGTGC CATGTTTCCT GAGAGACTGG GAGTTGCAGG	360
30	TGCACTTCAA AATCCATGGA CAAGGAAAGA AGAATCTGCA TGGGGATGGC TTGGCAATCT	420
	GGTACACAAG GAATCGGATG CAGCCAGGGC CTGTGTTTGG AAACATGGAC AAATTTGTGG	480
35	GGCTGGGAGT ATTTGTAGAC ACCTACCCCA ATGAGGAGAA GCAGCAAGAG CGGGTATTCC	540
	CCTACATCTC AGCCATGGTG AACAACGGCT CCCTCAGCTA TGATCATGAG CGGGATGGGC	600
40	GGCCTACAGA GCTGGGAGGC TGCACAGCCA TTGTCCGCAA TCTTCATTAC GACACCTTCC	660
40	TGGTGATTCG CTACGTCAAG AGGCATTTGA CGATAATGAT GGATATTGAT GGCAAGCATG	720
	AGTGGAGGGA CTGCATTGAA GTGCCCGGAG TCCGCCTGCC CCGCGGCTAC TACTTCGGCA	780
45	CCTCCTCCAT CACTGGGGAT CTCTCAGATA ATCATGATGT CATTTCCTTG AAGTTGTTTG	840
	AACTGACAGT GGAGAGAACC CCAGAAGAGG AAAAGCTCCA TCGAGATGTG TTCTTGCCCT	900
~~	CAGTGGACAA TATGAAGCTG CCTGAGATGA CAGCTCCACT GCCGCCCCTG AGTGGCCTGG	960
50	CCCTCTTCCT CATCGTCTT TTCTCCCTGG TGTTTTCTGT ATTTGCCATA GTCATTGGTA	1020
	TCATACTCTA CAACAAATGG CAGGAACAGA GCCGAAAGCG CTTCTACTGA GCCCTCCTGC	1080
55	TGCCACCACT TTTGTGACTG TCACCCATGA GGTATGGAAG GAGCAGGCAC TGGCCTGAGC	1140
	ATGCAGCCTG GAGAGTGTTC TTGTCTCTAG CAGCTGGTTG GGGACTATAT TCTGTCACTG	1200
60	GAGTTTTGAA TGCAGGGACC CCGCATTCCC ATGGTTGTGC ATGGGGACAT CTAACTCTGG	1260

	TCTGGGAAGC	CACCCACCCC	AGGGCAATGC	TGCTGTGATG	TGCCTTTCCC	TGCAGTCCTT	1320
	CCATGTGGGA	GCAGAGGTGT	GAAGAGAATT	TACGTGGTTG	TGATGCCAAA	ATCACAGAAC	1380
5	AGAATTTCAT	AGCCCAGGCT	GCCGTGTTGT	TTGACTCAGA	AGGCCCTTCT	ACTTCAGTTT	1440
	TGAATCCACA	AAGAATTAAA	AACTGGTAAC	ACCACAGGCT	TTCTGACCAT	CCATTCGTTG	1500
10	GGTTTTGCAT	TTGACCCAAC	CCTCTGCCTA	CCTGAGGAGC	TTTCTTTGGA	AACCAGGATG	1560
10	GAAACTTCTT	CCCTGCCTTA	CCTTCCTTTC	ACTCCATTCA	TIGTCCTCTC	TGTGTGCAAC	1620
	CTGAGCTGGG	AAAGGCATTT	GGATGCCTCT	CTGTTGGGGC	CTGGGGCTGC	AGAACACACC	1680
15	TGCGTTTCAC	TGGCCTTCAT	TAGGTGGCCC	TAGGGAGATG	GCTTTCTGCT	TTGGATCACT	1740
	GTTCCCTAGC	ATGGGTCTTG	GGTCTATTGG	CATGTCCATG	GCCTTCCCAA	TCAAGTCTCT	1800
20	TCAGGCCCTC	AGTGAAGTTT	GGCTAAAGGT	TGGTGTAAAA	ATCAAGAGAA	GCCTGGAAGA	1860
20	CATCATGGAT	GCCATGGATT	AGCTGTGCAA	CTGACCAGCT	CCAGGTTTGA	TCAAACCAAA	1920
	AGCAACATTT	GTCATGTGGT	CTGACCATGT	GGAGATGTTT	CTGGACTTGC	TAGAGCCTGC	1980
25	TTAGCTGCAT	GTTTTGTAGT	TACGATTTTT	GGAATCCCAC	TTTGAGTGCT	GAAAGTGTAA	2040
	GGAAGCTTTC	TTCTTACACC	TTGGGCTTGG	ATATTGCCCA	GAGAAGAAAT	TTGGCTTTTT	2100
30	TTTTCTTAAT	GGACAAGAGA	CAGTTGCTGT	TCTCATGTTC	CAAGTCTGAG	AGCAACAGAC	2160
50	CCTCATCATC	TGTGCCTGGA	AGAGTTCACT	GTCATTGAGC	AGCACAGCCT	GAGTGCTGGC	2220
	CTCTGTCAAC	CCTTATTCCA	CTGCCTTATT	TGACAAGGG	TTACATGCTG	CTCACCTTAC	2280
35	TGCCCTGGGA	TTAAATCAGT	TACAGGCCAG	AGTCTCCTTG	GAGGGCCTGG	AACTCTGAGT	2340
	CCTCCTATGA	ACCTCTGTAG	CCTAAATGAA	ATTCTTAAAA	TCACCGATGG	AACCAAAAAA	2400
40	АААААААА	AAAAAAAA	АААААААА	AA			2432
	(2) INFORMA	TION FOR SE	O TD NO: 51				
45		SEQUENCE CH					
		(A) LENG	GTH: 2340 ba	ase pairs			
50		(C) STRA	ANDEDNESS: O	double			
	(xi)	SEQUENCE I			. 51.		
	GACGCTGGGG					GCCCCCC	60
55	ATTAGTATGC						120
	AGGCAGCGGC						120
50	GAAGTGATTG						240
			· -				240

	GGCCCAGCTT GTTATTAAAA GACATCCTCA AATGTACATT GCTTGTGTTT GGAGTGTGGA	300
_	TCCTTTATAT CCTCAAGTTA AATTATACTA CTGAAGAATG TGACATGAAA AAAATGCATT	360
5	ATGTGGACCC TGACCATGTA AAGAGAGCTC AGAAATATGC TCAGCAAGTC TTGCAGAAGG	420
	AATGTCGTCC CAAGTTTGCC AAGACATCAA TGGCGCTGTT ATTTGAGCAC AGGTATAGCG	480
10	TGGACTTACT CCCTTTTGTG CAGAAGGSCC CCAAAGACAG TGAAGCTGAG TCCAAGTACG	540
	ATCCTCCTTT TGGGTTCCGG AAGTTCTCCA GTAAAGTCCA GACCCTCTTG GAACTCTTGC	600
15	CAGAGCACGA CCTCCCTGAA CACTTGAAAG CCAAGACCTG TCGGCGCTGT GTGGTTATTG	660
15	GAAGCGGAGG AATACTGCAC GGATTAGAAC TGGGCCACAC CCTGAACCAG TTCGATGTTG	720
	TGATAAGGTT AAACAGTGCA CCAGTTGAGG GATATTCAGA ACATGTTGGA AATAAAACTA	780
20	CTATAAGGAT GACTTATCCA GAGGGCGCAC CACTGTCTGA CCTTGAATAT TATTCCAATG	840
	ACTTATTTGT TGCTGTTTTA TTTAAGAGTG TTGATTTCAA CTGGCTTCAA GCAATGGTAA	900
25	AAAAGGAAAC CCTGCCATTC TGGGTACGAC TCTTCTTTTG GAAGCAGGTG GCAGAAAAAA	960
2,3	TCCCACTGCA GCCAAAACAT TTCAGGATTT TGAATCCAGT TATCATCAAA GAGACTGCCT	1020
	TTGRACATCC TTCAGTACTC AGAGCCTCAG TCAAGGTTCT GGGGGCCGAG ATAAGAACGT	1080
30	CCCCACAATC GGTGTCATTG CCGTTGTCTT AGCCACACAT CTGTGCGATG AAGTCAGTTT	1140
	GGCGGGTTTT GGATATGACC TCAATCAACC CAGAACACCT TTGCACTACT TCGACAGTCA	1200
35	ATGCATGGCT GCTATGAACT TTCAGACCAT GCATAATGTG ACAACGGAAA CCAAGTTCCT	1260
33	CTTAAAGCTG GTCAAAGAGG GAGTGGTGAA AGATCTCAGT GGAGGCATTG ATCGTGAATT	1320
	TTGAACACAG AAAACCTCAG TTGAAAATGC AACTCTAACT CTGAGAGCTG TTTTTGACAG	1380
40	CCTTCTTGAT GTATTTCTCC ATCCTGCAGA TACTTTGAAG TGCAGCTCAT GTTTTTAACT	1440
	TTTAATTTAA AAACACAAAA AAAATTTTAG CTCTTCCCAC TTTTTTTTTC CTATTTATTT	1500
45	GAGGTCAGTG TITGTTTTTG CACACCATTT TGTAAATGAA ACTTAAGAAT TGAATTGGAA	1560
73	AGACTTCTCA AAGAGAATTG TATGTAACGA TGTTGTWTTG ATTTTTAAGA AAGTAATTTA	1620
	ATTTGTAAAA CTTCTGCTCG TTTACACTGC ACATTGAATA CAGGTAACTA ATTGGAAGGA	1680
50	GAGGGGAGGT CACTCTTTTG ATGGTGGCCC TGAACCTCAT TCTGGTTCCC TGCTGCGCTG	1740
	CTTGGTGTGA CCCACGGAGG ATCCACTCCC AGGATGACGT GCTCCGTAGC TCTGCTGCTG	1800
55	ATACTGGGTC TGCGATGCAG CGGCGTGAGG CCTGGGCTGG TTGGAGAAGG TCACAACCCT	1860
<i>J J</i>	TCTCTGTTGG TCTGCCTTCT GCTGAAAGAC TCGAGAACCA ACCAGGGAAG CTGTCCTGGA	1920
	GGTCCCTGGT CGGAGAGGGA CATAGAATCT GTGACCTCTG ACAACTGTGA AGCCACCCTG	1980
60	GGCTACAGAA ACCACAGTCT TCCCAGCAAT TATTACAATT CTTGAATTCC TTGGGGATTT	2040

	TITACTGCCC TITCAAAGCA CTTAAGTGTT AGATCTAACG TGTTCCAGTG TCTGTCTGAG	2100
5	GTGACTTAAA AAATCAGAAC AAAACTTCTA TTATCCAGAG TCATGGGAGA GTACACCCTT	2160
	TCCAGGAATA ATGTTTTGGG AAACACTGAA ATGAAATCTT CCCAGTATTA TAAATTGTGT	2220
	ATTTAAAAAA AAGAAACTTT TCTGAATGCC TACTGGCGGT GTATACCAGG CAGTGTGCCA	2280
10	GTTTAAAAAG ATGAAAAAGA ATAAAAACTT TTGAGGAAMA AAAAAAAAAA AAAAACTCGA	2340
15	(2) INFORMATION FOR SEQ ID NO: 52:	
	(i) SEQUENCE CHARACTERISTICS:	
20	(A) LENGTH: 601 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 52:	
25	AGTAGGGGAG ACTGAGACTG ACCGGTAGCC AGGCAGGCGG ACGACGCACG CCCGGACAGA	60
	CTGAGCAGGC GCCGGAGAAC CACTCACAGG TTCCCCCCGC CTTTCCCTTT GAAANCTAGG	120
30	CTTTTGCCTT TCCCGTGGCG CCCGAGAGAG AATGCTGGAC TCTGCCGACT TCAGCGCAAC	180
30	TAANGATTTC TCAAGCTAGG GGACAAACGA TCAGCCCAAT CCTGAGAAGG GGGGAACCAA	240
	GCACCCCGTC CCCATCCCCC TCCCCTCCCC CGACTAAACT CGGGCGCCAA ACCCAGCCCT	300
35	TCTCTAACCA CCCTACTTCC TCCTCTCCTT TCTAGCATGG TGGCTGTATG GACAGTCTGA	360
	CAGAACAGAG ACTGACATCT CCCAATCTGC CGGCCCCCCA CCTGGAACAC TACAGTGTTC	420
40	TGCATTGCAC CATGACCCTG GATGTGCAAA CTGTAGTCGT TTTTGCCGTG ATTGTAGTCC	480
	TCCTGCTTGT CAATGTCATA CTCATGTTTT TCCTGGGAAC GCGCTGAATG GAGTCCAGNC	540
	ACCTGAGCTG TCGCGAACTC TCGCTTTGAT TTCATCCCGA GAGCCACCGA GAAGAAAAAA	600
45	A	601
50	(2) INFORMATION FOR SEQ ID NO: 53:	
	(i) SEQUENCE CHARACTERISTICS:	
55	(A) LENGTH: 359 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double	
-	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 53:	
60	CTCGTGCCGA ATTCGGCACG AGAGATGGTA CTTTTAAGAG GTAATTAGGT TGCTAAGATG	60

	GATTAACATC TTTCTCTTGA CACTGAGACT GGGTTCTCCT GGGAATGGTT AGTTCCCAAG	120
_	AGAGTGAGTT GTTATAAAAC AATGCTGCCT CTTCTATTTT GCGCTTTTTG TTTGCACAAA	180
5	CTCGGTCCCC TTCTGTTTCT CTACGATGTT TTGATGCRGC ATGAGGCAGT CATGAGAACC	240
	CACCAGATAC AGCTGCCTGA TCCTGAATTT CCCAGCCAAC AGAACCAAGT GCTAAATAAA	300
10	ACTCTTTTTA ATAAGTTAAA AAAAAAAAA AAAAAAAAA AANAAANANA AAAAAA	359
. ~		
15	(2) INFORMATION FOR SEQ ID NO: 54:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1141 base pairs	
	(B) TYPE: nucleic acid	
20	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 54:	
25	GGCACGAGCT GCTGAGGCGT GAGAATGGCG TCCCGCGGCC GGCGTCCGGA GCATGGCGGA	60
	CCCCCAGAGC TGTTTTATGA CGAGACAGAA GCCCGGAAAT ACGTTCGCAA CTCACGGATG	120
20	ATTGATATCC AGACCAGGAT GGCTGGGCGA GCATTGGAGC TTCTTTATCT GCCAGAGAAT	180
30	AAGCCCTGTT ACCTGCTGGA TATTGGCTGT GGCACTGGGC TGAGTGGAAG TTATCTGTCA	240
	GATGAAGGGC ACTATTGGGT GGGCCTGGAT ATCAGCCCTG CCATGCTGGA TGAGGCTGTG	300
35	GACCGAGAGA TAGAGGGAGA CCTGCTGCTG GGGGATATGG GCCAGGGCAT CCCATTCAAG	360
	CCAGGCACAT TTGATGGTTG CATCAGCATT TCTGCTGTGC AGTGGCTCTG TAATGCTAAC	420
40	AAGAAGTCTG AAAACCCTGC CAAGCGCCTG TACTGCTTTT TTGCTTCTCT TTTTTCTGTT	480
40	CTCGTCCGGG GATCCCGAGC TGTCCTGCAG CTGTACCCTG AGAACTCAGA GCAGTTGGAG	540
	CTGATCACAA CCCAGGCCAC AAAGGCAGGC TTCTCCGGTG GCATGGTGGT AGACTACCCT	600
45	AACAGTGCCA AAGCAAAGAA ATTCTACCTC TGCTTGTTTT CTGGGCCTTC GACCTTTATA	660
	CCAGAGGGC TGAGTGAAAA TCAGGATGAA GTTGAACCCA GGGAGTCTGT GTTCACCAAT	720
50	GAGAGGTTCC CATTAAGGAT GTCGAGGCGG GGAATGGTGA GGAAGAGTCG GGCATGGGTG	780
50	CTGGAGAAGA AGGAGCGGCA CAGGCGCCAG GGCAGGGAAG TCAGACCTGA CACCCAGTAC	840
	ACCGGCCGCA AGCGCAAGCC CCGCTTCTAA GTCACCACGC GGTTCTGGAA AGGCACTTGC	900
55	CTCTGCACTT TTCTATATTG TTCAGCTGAC AAAGTAGTAT TTTAGAAAAG TTCTAAAGTT	960
	ATAAAAATGT TTTCTGCAGT AAAAAAAAAG TTCTCTGGGC CGGGCGTGGT GGCTCACACC	1020
	TGTAATCCCA GCACCTTGGG AGGCTGAGGT GGGAGGATCA TTTGAGGCCA GGAGTTTGAG	1080

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	ACCTGCCTGG GCAACATAAT GAAACTTCCT TTCCAGGGAG AAAAAAAAAA	1140
	A	1141
5		
	(2) TITONATION DO TO TO TO	
10	(2) INFORMATION FOR SEQ ID NO: 55:	
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 1560 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 55:	
	TCCTTCTCTG GGGCGGTCGC GTTGGCAGCG GATGCGGGAA GCCGGACTCT GGGCGTCATG	60
20	TACTACAAGT TTAGTGGCTT CACGCAGAAG TTGGCAGGAG CATGGGCTTC GGAGGCCTAT	120
	AGCCCGCAGA TINAAAGCCT GTGGTTTCCA CAGAAGCACC ACCTATCATA TTTGCCACAC	180
25	CAACTAAACT GACCTCCGAT TCCACAGTGT ATGATTATGC TGGGAAAAAC AAAGTTCCAG	240
	AGCTACAAAA GTTTTTCCAG AAAGCTGATG GTGTGCCCGT CTACCTGAAA CGAGGCCTGC	300
	CTGACCAAAT GCTTTACCGG ACCACCATGG CGCTGACTGT GGGAGGGACC ATCTACTGCC	360
30	TGATCGCCCT CTACATGGCT TCGCAGCCCA AAAACAAATG AGTTAGGCTG CAGAGGACTG	420
	GTTTGTTTTT TGGCATAAAC CCTTTGAAGT TCCTTTTTCA TTGTTAAATT AAAATTTTTT	480
35	TTTTTACTTG GATGGCTTAA CATTTTTGCA AGAAAAATAG GAAGATATGA AGATGATGTT	540
	TTGGTTTGTT TATGAAATGC ATATGGCTTG TCAGAGCTCA TTCGACAGTT AAAGCCATTG	600
	TTTAAAGAAA CGGTGCTTTG CTCTGTGTTT GTGCTCCTGA TTTCCCTGGA GGTTCTGGAT	660
40	GAAGGCTGAA CACAGGCTTG TTAATGTCAG TCTGTGCTGA GGACCTCAGG GACTTGAGGT	720
	TGCATTTTTG AGCATGGGGT GCAGGAGCCT TTCTGGATTT GGATGTGGCT ATGGAAAGAA	780
45	CACAGAAGCC AAGGTCATGT GCATGAAATG AGGAGTTTGA GTTAGTCACC TCGGGGATTT	840
	TTTCCATTTT GCAGTAAAAT GTTAAATTAA TGTAGCCTGC CTCTATTTGT TGGGCAGGTA	900
	ATTTCAAAGG GTTATTTGCC TCATCTCCTA TCTTTAGTGA AATCTTATGT GTAATTGTGT	960
50	GTATTTATTC CACCGTGGGA ACAGAGAATA CCTGTTTAGT GTTGCACTTT AGACTGGTGT	1020
	CTGTTTTGTT AATGCAGCTG TGCCACAAAT TCTCCTTTAT CTTTTAAAAA TGTTATAGCT	1080
55	TTAAATTTTG ATTTATTTTG ACTGTGGAAT AAATACATGA ATGAAAAATT TTAAGTTTGA	1140
	AGTTCTTTGA ATGACCTTTC AGAGTAATTT CAGAACACCA GCAGCATCTT AAACCTGAGT	1200
	CTAATTTCTT TCTTGTTAAT TAGGCACCAG ATAATCTTTA TAAAATGGTC TTAAAAGCTA	1260
60	GTAATAGGAG CTTAATGGCA ATKGATGATT ACCACAKGGT TTTTTATAAA AACCTGCCTG	1320

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	CCCCTWAGTG AAAGGTACCT GTAACYCACA GTYCATTTAG ACACTAATTT CCTYTGCYGT	1380
_	CATGATTGGK AGACTTCACT TACCCTATAT TAATTTTGAA AAAAGGTGGA ATTTTATTAT	1440
5	ATATGAAGGA ATAGTTTGTA TCTTACCATA GCACAGAACA GTGACCTCTT GCTCAGGATA	1500
	AGATGTGGTG ATTTGAAAAT ACTCATAGTA GCCTTGCAGT GATACCTCTC TCNCTCTCTC	1560
10		
	(2) INFORMATION FOR SEQ ID NO: 56:	
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1507 base pairs	
	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
20	(D) TOPOLOGY: linear	
_ 0	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 56:	
	GGAACGCAGA GCGGAGCGTG GAGAGCGGAG CGAAGCTGGA TAACAGGGGA CCGATGATGT	60
25	GGCGACCATC AGTTCTGCTG CTTCTGTTGC TACTGAGGCA CGGGGCCCAG GGGAAGCCAT	120
	CCCCAGACGC AGGCCCTCAT GGCCAGGGGA GGGTGCACCA GGCGGCCCCC CTGAGCGACG	180
20	CTCCCCATGA TGACGCCCAC GGGAACTTCC AGTACGACCA TGAGGCTTTC CTGGGACGGG	240
30	AAGTGGCCAA GGAATTCGAC CAACTCACCC CAGAGGAAAG CCAGGCCCGT CTGGGGCGGA	300
	TCGTGGACCG CATGGACCGC GCGGGGGACG GCGACGCTG GGTGTCGCTG GCCGAGCTTC	360
35	GCGCGTGGAT CGCGCACACG CAGCAGCGGC ACATACGGGA CTCGGTGAGC GCGGCCTGGG	420
	ACACGTACGA CACGGACCGC GACGGGCGTG TGGGTTGGGA GGAGCTGCGC AACGCCACCT	480
40	ATGGCCACTA CGCGCCCGGT GAAGAATTTC ATGACGTGGA GGATGCAGAG ACCTACAAAA	540
40	AGATGCTGGC TCGGGACGAG CGGCGTTTCC GGGTGGCCGA CCAGGATGGG GACTCGATGG	600
	CCACTCGAGA GGAGCTGACA GCCTTCCTGC ACCCCGAGGA GTTCCCTCAC ATGCGGGACA	660
45	TCGTGATTGC TGAAACCCTG GAGGACCTGG ACAGAAACAA AGATGGCTAT GTCCAGGTGG	720
	AGGAGTACAT CGCGGATCTG TACTCAGCCG AGCCTGGGGA GGAGGAGCCG GCGTGGGTGC	780
50	AGACGGAGAG GCAGCAGTTC CGGGACTTCC GGGATCTGAA CAAGGATGGG CACCTGGATG	840
50	GGAGTGAGGT GGGCCACTGG GTGCTGCCCC CTGCCCAGGA CCAGCCCCTG GTGGAAGCCA	900
	ACCACCTGCT GCACGARAGC GACACGGACA AGGAYGGGCG GCTGAGCAAA GCGSAAATCC	960
55	TGGGTAATTG GAACATGTTT GTGGGCAGTC AGGCCACCAA CTATGGYGAG GACCTGACCC	1020
	GGCACCACGA TGAGCTGTGA GCMCCGNGCA CCTGCCACAG CCTCAGAGGC CCGCACAATG	1080
	ACCGGAGGAG GGGCCGCTGT GGTCTGGCCC CCTCCCTGTC CAGGCCCCGC AGGAGGCAGA	1140

	TGCAGTCCCA GGCATCCTCC TKCCCCTGGG CTCTCAGGGA CCCCCTGGGT CGGCTTCTGT	1200
	CCCTGTCACA CCCCCAACCC CAGGGAGGGG CTGTCATAGT CCCAGAGGAT AAGCAATACC	1260
5	TATTICTGAC TGAGTCTCCC AGCCCAGACC CAGGGACCCT NGGCCCCAAG CTCAGCTCTA	1320
	AGAACCGCCC CAACCCCTCC AGCTCCAAAT CTGAGCCTCC ACCACATAGA CTGAAACTCC	1380
10	CCTGGCCCCA GCCCTCTCCT GCCTGGCCTG GCCTGGGACA CCTCCTCTCT GCCAGGAGGC	1440
	AATAAAAGCC AGCGCCGGGA AAAAAAAAAA AAAAAAAAA AAAAAAAA	1500
	NAAAAAA	1507
15		
	(2) INFORMATION FOR SEQ ID NO: 57:	
20	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 450 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 57:	
	GAATTCGGCA CGAGCAGTGT CCAACACTGT AGCTGGTGCC TGCCAGGTTC CCAGTGGCTG	60
30	GGGTCACCAG GTCTGAAGAG AGATGTGCTG GCTGCGGGCA TGGGSCCAGA TCYTCCTGCC	120
	AGTITICYTC TCCYTCTTTC TCATCCAATT GCTTATCAGC TTCTCAGAGA ATGGTTTTAT	180
35	CCACAGCCCC AGGAACAATC AGAAACCAAG AGATGGGAAT RAAGAGGAAT GTGCTGTAAA	240
33	GAAGAGTTGT CAATTGTGCA CAGAAGATAA GAAATATATG ATGAATAGAT AATTGAAAAG	300
	AGATCCTCCA GAAAGAGCAG AAGGAAGTTT CTTCAATGGC TTCCTTCAGG ATTTTAATCA	360
40	TCCTTACAGC CTCTTTGAGA ATGATTGAAC TTCCAAATTC CCTGAAGTTA AAATTTTAAA	420
	TTCTATTAAA CATTTTTTCG AGTAAAAAA	450
45		
	(2) INFORMATION FOR SEQ ID NO: 58:	
50	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1147 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
55	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 58:	
	GGCACGAGAC CCATTGAGCA GAAGGAGGCC AGGTGGGAAA GCTCCTGGGA AGAGCAGCCA	60
60	GACTGGACAC TGGGCTGCTT GAGTCCTGAG TCACAATTCA GAATTCCTGG GCTCCCTGGG	120

	TGCATTCTAT CATTCCAGTT GAAAGTTTGC TTCCTTCCAG TCATGTGGCT CTTCATTCTA	180
	CTCTCCTTGG CTCTCATTTC AGATGCCATG GTCATGGATG AAAAGGTCAA GAGAAGCTTT	240
5	GTGCTGGACA CGGCTTCTGC CATCTGCAAC TACAATGCCC ACTACAAGAA TCACCCCAAA	300
	TACTGGTGCC GAGGCTATTT CCGTGACTAC TGCAACATCA TCGCCTTCTC CCCTAACAGC	360
10	ACCAATCATG TGGCCCTGAA GGACACAGGG AACCAGCTCA TTGTCACTAT GTCCTGCCTG	420
10	AACAAAGAAG ACACGGGCTG GTACTGGTGT GGCATCCAGC GGGACTTTGC CAGGGATGAC	480
	ATGGATTTTA CAGAGCTGAT TGTAACTGAC GACAAAGGAA CCTGGCCAAT GACTTTGGTC	540
15	TGGGAAAGAC TATCAGGCAC AAAACCAGAA GCTGCAAGGC TCCCAAAGTT GTCCGCAAGG	600
	CTGACCGCTC CAGGACGTCC ATTCTCATCA TTTGCATACT GATCACGGGT TTGGGAATCA	660
20	TCTCTGTAAT CAGTCATTTG ACCAAAAGGA GGAGAAGTCA AAGGAATAGA AGGGTAGGCA	720
20	ACACTTTGAA GCCCTTCTCG CGTGTCCTGA CTCCAAAGGA AATGGCTCCT ACTGAACAGA	780
	TGTGACTGAA GATTTTTTTA ATTTAGTTCA TAAAGTGATG CTACAACAGA ATAATCACCA	840
25	TGACAACTGG CCCCACACCT CAGAGACTGA TTCTGATCTC CCAGGAATTC TGAAGGTCCC	900
	TCTATCCTTG ACAACAATCA TTTGCAGCCA GGTAGCAACG GCAGTAGTCA GAGGAGCTAT	960
20	GATAGACCAC ACCCAAGCAA GGCTGCCCTC AAATAACATC TCAAGATCTT AGTTCTTATG	1020
30	CATTCCATCA GTCAGAAGTG AAGAAGAGGT GGAGAATCTG GATTGGGGAC CAGGAAATCA	1080
	CTTGTATTTT GTTAGCCAAT AAATTCCTAG CCAGTGTTGA ATGAAAAAAA AAAAAAAAAA	1140
35	даааааа	1147

40 (2) INFORMATION FOR SEQ ID NO: 59:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 777 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 59:

GCAGAGGCT CCTCAGAAGG GCGTGGCTC TCCAGTCTC CACAGTCCCC ACCATGCCCT 60

GTTGCCTTAC CGCTGACGTA GCTCACCCAT CTTTTACTTG CCTGGCTAAG ATGCATGGCA 120

TYWCATTTCC TCCTTGTTGC ACTGCAGTCA GTCCCTCACT GCCCCCATCT CCTGGAAGAG 180

GAGCATAAGC TTTGCAAGGT CAGCCACTTC TCTGGGGTCA CACTAGTTAC ATCAAGACAG 240

GACTCCAGCT CATATGTGCC AGTGCAGACA CTCTTCATCC ACCTGGGCC CTGGGCTTGG 300

60 GACCTGGYTC CTTGCACAGC AGARGACCCG GAGGCTGAGA GGAGCTTGCG GTTGTGTCAT 360

	AGTCACCTGG CCAGARGGAA CGTGAGCCCC TCCCAAGCTG CAGARGGARG GARCARGCGT	420
5	GGCTGTCAGC ACCGAGGTAG CAGAGAATTA ACATTCTTGT CAGCAGAGAA TGAAGCAGGA	480
	ATATAATTAA AACTTTGCCC TTGGAATAGC TGATTCATTT GAATTTTATT CCACACGTTT	540
	GAAAGAGGAA AGAAAATGTG AAGACTTGCA GCCTGGTTCT CGCCTGGCCT GGGCTGGCCC	600
10	AGCTGTCAGG CCCGGTTCCT TTCTGAGCAT TCAGTCCACT GATGTTGACT GAGGGCCAGG	660
	AGAGACCCTC AGCAGGGTAT TACCATATCA GCCTCCTATC GCTGCTGGGA GAAATTACCA	720
15	TGAATTCAGT GGCTTAAAAC AACACACGAG CCTCTCTGAG CCTACCCTGG CTCAGGA	777
13		
20	(2) INFORMATION FOR SEQ ID NO: 60: (i) SEQUENCE CHARACTERISTICS:	
25	(A) LENGTH: 1191 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 60:	
30	AAGANIGATT TTCCTTACTC TCCAAAGCGT CAGCATTTTG AAGTTTCTTT TATGAAAGTG	60
	GGGGCAAGAA TCAGGGTGAA AATGAGTGTA AACAAAGCCC ATCCTGTGGT CAGCACCCAC	120
	TGGAGGTGGC CAGCAGAGTG GCCTCAGATG TTCCTGCACC TGGCCCAGGA GCCCAGGACA	180
35	GAGGTCAAAT CTAGGCCCCT TGGTCTGGCT GGATTCATCA GGCAAGATTC GAAAACAAGA	240
	AAACCTCTAG AACAAGAAAC AATCATGTCT GCAGCAGATA CGGCACTGTG GCCCTATGGC	300
40	CATGGCAATC GTGAGCACCA AGAGAATGAG TTACAGAAAT ATCTCCAATA CAAAGACATG	360
	CATCTCCTGG ACAGTGGACA GTCGCTGGGA CACACACAC CACTTCAAGG CTCACACAAC	420
	CTAACAGCCT TAAATATCTG AAGAAACAGA ATCACGACAT TAAGTCAGCA GAGGGAGAGG	480
45	TAGGCTGAAG CAGCAGGAGG CCAATTTTAT ATCCCACAGA TTTTTTTAAA AATGACTCCC	540
	CAGCAAGGGG TGGGGAGAAA GCCACTGATT TAGGAGAGTT CTTGGCTCAG CCAACCACTG	600
50	CGGTTATCTA CACGTTTTAC AAAGGCACRG AAGTAGAGAG GGGCTGCACT CACGACCCTC	660
- 0	CCCAGGCCC GCACAGCCAG ACACGGTGGG TTCTTCCTTT TTCCCTTCTG GCCTTGGTGG	720
	AATTCCTACC ACGGTGGCCT CTGCCTTTGG GACAATGCCT TCATGCTCAT CCCCGGGTCA	780
55	AGGATGGAGT CTGTTACCAT TTTCCAGGGG AAATTCCAAG GACCAGCCCC GCCTCATTAC	840
	GTTCACCCCA CAGGAAGGTG ATCTGGAAAG CCTGTAAACA CGTACTCTGG GTGGCTGAGT	900
60	GGTGTCACCA AGCTGCTTTT GTGCAGGGCT GAAGCACAGA CAAGAGGGCA GGCAGCTGCC	960

1200

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	GGAGGCCTGA AGTGGGGAGA GATCCCCGCA GGCCTGCAGG AGCCAGGGAG AACCTCCAAC	1020
	TGGATCTAAA CTGTGGGACA GCCCAGGCGT GCCCCTCTTC ACATGGCTCC CAGGCTCCCT	1080
5	CAAAGCCCTT CCCAGGCCCT GCAGGAAGAG AGGGAGGGTG AGGAGAGGCA GGGAGGGCAG	1140
	AGGTCGCCTG AAAGCCTGGG CTCCGAACTC CCTCAGCAGA GCTTTAAAGT G	1191
10		
	(2) INFORMATION FOR SEQ ID NO: 61:	
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 1580 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 61:	
	CCCCGCCCC CGCCCACGAA GGAAGTGGCT GCTGCTCCGG CGCGGACCCA GAGCCGGTTC	60
2.5	GGCGCGTCGA CTGCCCAGAG TCCGCGGCCG GGCGCGGGAG GAGCCAAGCC GCCATGGCCT	120
25	ACCACAGCTT CCTGGTGGAG CCCATCAGCT GCCACGCCTG GAACAAGGAC CGCACCCAGA	180
	TIGCCATCTG CCCCAACAAC CATGAGGTGC ATATCTATGA AAAGAGCGGT GCCAAATGGA	240
30	CCAAGGTGCA CGAGCTCAAG GAGCACAACG GGCAGGTGAC AGGCATCGAC TGGGCCCCCG	300
	AGAGTAACCG TATTGTGACC TGCGGCACAG ACCGCAACGC CTACGTGTGG ACGCTGAAGG	360
	GCCGCACATG GAAGCCCACG CTGGTCATCC TGCGGATCAA CCGGGCTGCC CGCTGCGTGC	420
35	GCTGGGCCCC CAACGAGAAC AAGTTTGCTG TGGGCAGCGG CTCTCGTGTG ATCTCCATCT	480
	GTTATTTCGA GCAGGAGAAT GACTGGTGGG TTTGCAAGCA CATCAAGAAG CCCATCCGCT	540
40	CCACCGTCCT CAGCCTGGAC TGGCACCCCA ACAATGTGCT GCTGGCTGCC GGCTCCTGTG	600
	ACTTCAAGTG TCGGATCTTT TCAGCCTACA TCAAGGAGGT GGAGGAACGG CCGGCACCCA	660
45	CCCCGTGGGG CTCCAAGATG CCCTTTGGGG AACTGATGTT CGAATCCAGC AGTAGCTGCG	720
45	GCTGGGTACA TGGCGTCTGT TTCTCAGCCA GCGGGAGCCG CGTGGCCTGG GTAAGCCACG	780
	ACAGCACCGT CTGCCTGGCT GATGCCGACA AGAAGATGGC CGTCGCGACT CTGGCCTCTG	840
50	AAACACTACC ACTGCTGGCG CTGACCTTCA TCACAGACAA CAGCCTGGTG GCAGCGGCC	900
	ACGACTGCTT CCCGGTGCTG TTCACCTATG ACGCCGCCGC GGGGATGCTG AGCTTCGGCG	960
55	GGCGGCTGGA CGTTCCTAAG CAGAGCTCGC AGCGTGGCTT GACGGCCCGC GAGCGCTTCC	1020
	AGAACCTGGA CAAGAAGGCG AGCTCCGAGG GTGGCACGGC TGCGGGCGCG GGCCTAGACT	1080

CGCTGCACAA GAACAGCGTC AGCCAGATCT CGGTGCTCAG CGGCGGCAAG GCCAAGTGCT

CGCAGTTCTG CACCACTGGC ATGGATGGCG GCATGAGTAT CTGGGATGTG AAGAGCTTGG

	AGTCAGCCTT GAAGGACCTC AAGATCAAAT GACCTGTGAG GAATATGTTG CCTTCATCCT	1260
5	AGCTGCTGGG GAAGCGGGA GAGGGGTCAG GGAGGCTAAT GGTTGCTTTG CTGAATGTTT	1320
	CTGGGGTACC AATACGAGTT CCCATAGGGG CTGCTCCCTC AAAAAGGGAG GGGACAGATG	1380
	GGGAGCTTTT CTTACCTATT CAAGGAATAC GTGCCTTTTT CTTAAATGCT TTCATTTATT	1440
10	GAAAAAAAA AAAAATGCCC CCAAAGCACT ATGCTGGTCA TGAACTGCTT CAAAATGTGG	1500
	AGGTAATAAA ATGCAACTGT GTAAAAAAAA AAAAAAAAA AAATGACCCT CGCGATCTAG	1560
15	AACTAGNCGG ACGCNTGGGT	1580
20	(2) INFORMATION FOR SEQ ID NO: 62:	
25	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1117 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 62:	
	GGCACGAGGC GCGATGCAGC ACAGGCTAGA GGCTGCGCAA SGCGGGGGCC CGCCCC'IGGG	60
30	ACCCTCCGGG CCGGGCGGTT TGGCCCCTTA GCGCCCGGGC GTCGGGGCGG TAAAAGGCCG	120
	GCAGAAGGGA GGCACTTGAG AAATGTCTTT CCTCCAGGAC CCAAGTTTCT TCACCATGGG	180
35	GATGTGGTCC ATTGGTGCAG GAGCCCTGGG GGCTGCTGCC TTGGCATTGC TGCTTGCCAA	240
	CACAGACGTG TTTCTGTCCA AGCCCCAGAA AGCGGCCCTG GAGTACCTGG AGGATATAGA	300
40	CCTGAAAACA CTGGAGAAGG AACCAAGGAC TTTCAAAGCA AAGGAGCTAT GGGAAAAAAA	360
40	TGGAGCTGTG ATTATGGCCG TGCGGAGGCC AGGCTGTTTC CTCTGTCGAG AGGAAGCTGC	420
	GGATCTGTCC TCCCTGAAAA GCATGTTGGA CCAGCTGGGC GTCCCCCTCT ATGCAGTGGT	480
45	AAAGGAGCAC ATCAGGACTG AAGTGAAGGA TTTCCAGCCT TATTTCAAAG GAGAAATCTT	540
	CCTGGATGAA AAGAAAAAGT TCTATGGTCC ACAAAGGCGG AAGATGATGT TTATGGGATT	600
50	TATCCGTCTG GGAGTGTGGT ACAACTTCTT CCGAGCCTGG AACGGAGGCT TCTCTGGAAA	660
50	CCTGGAAGGA GAAGGCTTCA TCCTTGGGGG AGTTTTCGTG GTGGGATCAG GAAAGCAGGG	720
	CATTCTTCTT GAGCACCGAG AAAAAGAATT TGGAGACAAA GTAAACCTAC TTTCTGTTCT	780
55	GGAAGCTGCT AAGATGATCA AACCACAGAC TTTGGCCTCA GAGAAAAAAT GATTGTGTGA	840
	AACTGCCCAG CTCAGGGATA ACCAGGGACA TTCACCTGTG TTCATGGGAT GTATTGTTTC	900
60	CACTCGTGTC CCTAAGGAGT GAGAAACCCA TTTATACTCT ACTCTCAGTA TGGATTATTA	960

	ATGTATTTTA ATATTCTGTT TAGGCCCACT AAGGCAAAAT AGCCCCAAAA CAAGACTGAC	1020
	AAAAATCTGA AAAACTAATG AGGATTATTA AGCTAAAACC TGGGAAATAG GAGGCTTWAA	1080
5	ATGACTGCCM GCTGGTGCRT GCTCACACTT GGCCCAC	1117
10	(2) INFORMATION FOR SEQ ID NO: 63:	
10		
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 361 base pairs	
15	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
13	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 63:	
20	CCCACGCGTG CKGGCGCCTG GCAGCCACCG CCTGGGAGGT TACTGTAAGG CCCGCAGCTC	60
	CCGCCAGCTC CCGCGGACTS CTGCCGCCTC CTTACCATGA AGCCAGTAAG TCGTCGCACG	120
25	CTGGACTGGA TTTATTCAGT GTTGCTGCTT GCCATCGTTT TAATCTCCTG GGGCTGCATC	180
40	ATCTATGCTT CGATGGTGTC TGCAAGACGA CAGCTAAGGA AGAAATACCC AGACAAAATC	240
	TTTGGGACGA ATGAAAATTT GTAACTCTTC TGGATTTAAT TATCTGAAAA TACAGTTCTT	300
30	TCCCTCATGC TTATGTAGAT ATAAAAATAA AATTCATAAT GCAAAAAAAA AAAAAAAAAA	360
	G	361
35		
55	(2) INFORMATION FOR SEQ ID NO: 64:	
40	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1668 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
45	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 64:	
	GGCACGAGGT CTGCCAAGCT ATAGACCATG GCTGTGAACA CATTTGTGTG AACAGTGACG	60
50	ACTCATACAC GTGCGAGTGC TTGGAGGGAT TCCGGCTCGC TGAGGATGGG AAACGCTGCC	120
50	GAAGAAGGAT GTCTGCAAAT CAACCCACCA TGGCTGCGAA CACATTTGTG TTAATAATGG	180
	GAATTCCTAC ATCTGCAAAT GCTCAKAGGG ATTTGTTCTA GCTGAGGACG GAAGACGGTG	240
55	CAAGAAATGC ACTGAAGGCC CAATTGACCT GGTCTTTGTG ATCGATGGAT CCAAGAGTCT	300
	TGGAGAAGAG AATTTTGAGG TCGTGAAGCA GTTTGTCACT GGAATTATAG ATTCCTTGAC	360
60	AATTTCCCCC AAAGCCGCTC GAGTGGGGCT GCTCCAGTAT TCCACACAGG TCCACACAGA	420
OU		

ATACATGGGA AAGGGCTCTA TGACTGGGCT GGCCCTGAAA CACATGTTTG AGAGA 5 TACCCAAGGA GAAGGGGCCA GGCCCTTTCC ACAAGGGTGC CCAGAGCAGC CATTC ACCGACGGAC GGGCTCAGGA TGACGTCTCC GAGTGGGCCA GTAAAGCCAA GGCCA	
CATTLE REALDESTANCE CANADAGE CATTLE	FTGTTC 600
ACCGACGGAC GGGCTCAGGA TGACGTCTCC GAGTGGGCCA GTAAAGCCAA GGCCA	
	AATGGT 660
ATCACTATGT ATGCTGTTGG GGTAGGAAAA GCCATTGAGG AGGAACTACA AGAGA	ATTGCC 720
TCTGAGCCCA CAAACAAGCA TCTCTTCTAT GCCGAAGACT TCAGCACAAT GGATG	GAGATA 780
AGTGAAAAAC TCAAGAAAGG CATCTGTGAA GCTCTAGAAG ACTCCGATGG AAGAC	CAGGAC 840
15 TCTCCAGCAG GGGAACTGCC AAAAACGGTC CAACAGCCAA CAGTGCAACA CAGAT	PATCTG 900
TTTGAAGAAG ACAATCTTTT ACGGTCTACA CAAAAGCTTT CCCATTCAAC AAAAC	CCTTCA 960
GGAAGCCCTT TGGAAGAAAA ACACGATCAA TGCAAATGTG AAAACCTTAT AATGT	TTCCAG 1020
AACCTTGCAA ACGAAGAAGT AAGAAAATTA ACACAGCGCT TAGAAGAAAT GACAC	CAGAGA 1080
ATGGAAGCCC TGGAAAATCG CCTGAGATAC AGATGAAGAT TAGAAATCGC GACAC	CATTTG 1140
25 TAGTCATTGT ATCACGGATT ACAATGAACG CAGTGCAGAG CCCCAAAGCT CAGGC	CTATTG 1200
TTAAATCAAT AATGTTGTGA AGTAAAACAA TCAGTACTGA GAAACCTGGT TTGCC	CACAGA 1260
ACAAAGACAA GAAGTATACA CTAACTTGTA TAAATTTATC TAGGAAAAAA ATCCT	TCAGA 1320
ATTCTAAGAT GAATTTACCA GGTGAGAATG AATAAGCTAT GCAAGGTATT TTGTA	ATATA 1380
CTGTGGACAC AACTTGCTTC TGCCTCATCC TGCCTTAGTG TGCAATCTCA TTTGA	ACTATA 1440
35 CGATAAAGTT TGCACAGTCT TACTTCTGTA GAACACTGGC CATAGGAAAT GCTGT	TTTTT 1500
TGTAYTGGAC TTTACCTTGA TATATGTATA TGGATGTATG CATAAAATCA TAGGA	CATAT 1560
GTACTTGTGG AACAAGTTGG ATTTTTTATA CAATATTAAA ATTCACCACT TCAGA	GRAAA 1620
AAAAAAAA AAAAAAAAA AAAAAAAA AAAAAAAA AAAA	1668
45 (2) INFORMATION FOR SEQ ID NO: 65:	
(i) SEQUENCE CHARACTERISTICS:	
(A) LENGTH: 1353 base pairs (B) TYPE: nucleic acid	
(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 65:	
55	CGTTG 60
GGGTCGACCC ACGCGTCCGC CCACGCGTCC GGATGGCTGC GCTGTTCCTTC ACACAC	
GGGTCGACCC ACGCGTCCGC CCACGCGTCC GGATGGCTGC GCTGTTGCTG AGACA GTCGTCATTG CCTCCGAGCC CACTTTAGCC CTCAGCTCTG TATCAGAAAT GCTGT	TCCTT 120

	GTCCTCTGTC TCCCCACATT ACTATCTACA GTTGGTCTCT TCCCATGGCG ATGTCCATCT	240
_	GCCACCGTGG CACTGGTATT GCTTTGAGTG CAGGGGTCTC TCTTTTTGGC ATGTCGGCCC	300
5	TGTTACTCCC TGGGAACTTT GAGTCTTATT TGGAACTTGT GAAGTCCCTG TGTCTGGGGC	360
	CAGCACTGAT CCACACAGCT AAGTTTGCAC TTGTCTTCCC TCTCATGTAT CATACCTGGA	420
10	ATGGGATCCG ACACTTGATG TGGGACCTAG GAAAAGGCCT GAAGATTCCC CAGCTATACC	480
	AGTCTGGAGT GGTTGTCCTG GTTCTTACTG TGTTGTCCTC TATGGGGCTG GCAGCCATGT	540
1.5	GAAGAAAGGA GGCTCCCAGC ATCATCTTCC TACACATTAT TACATTCACC CATCTTTCTG	600
15	TTTGTCATTC TTATCTCCAG CCTGGGAAAA GTTCTCCTTA TTTGTTTAGA TCCTTTTGTA	660
	TTTTCAGATC TCCTTGGAGC AGTAGAGTAC CTGGTAGACC ATAATAGTGG AAAAGGGTCT	720
20	AGTTTTCCCC TTGTTTCTAA AGATGAGGTG GCTGCAAAAA CTCCCCTTTT TTGCCCACAG	780
	CTTGCCTACT CTCGGCCTAG AAGCAGTTAT TCTCTCTCCA TATTGGGCTT TGATTTGTGC	840
25	TGAGGGTCAG CTTTTGGCTC CTTCTTCCTG AGACAGTGGA AACAATGCCA GCTCTGTGGC	900
25	TICTGCCCTG GGGATGGGCC GGGTTGGGG GTGGGTTGGT GAGGCTTTGG GTGCCACTGC	960
	CTGTGGGTTG CTGGCTTAAA GGACAATTCT CTTCATTGGT GAGAGCCCAG GCCATTAACA	1020
30	CCTACACAGT GTTATTGAAA GAAGAGAGGT GGGGGTGGAG GGGAATTAGT CTGTCCCAGC	1080
	TAGAGGGAGA TAAAGAGGGC TAGTTAGTTC TTGGAGCAGC TGCTTTTGAG GAGAAAATAT	1140
35	ATAGCTTTGG ACACGAGGAA GATCTAGAAA ATTATCATTG AACATATTAA TGGTTATTTC	1200
33	TTTTTCTTGG ATTTCCAGAA AAGCCTCTTA ATTTTATGCT TTCTCATCGA AGTAATGTAC	1260
	CCTTTTTTC TGAAACTGAA TTAAATACTC ATTTTATCTT TGAAAAAAAA AAAAAAAACC	1320
40	TNGGGGGGG CCCCGGACCC NAATTGGCCC TAT	1353
45	(2) INFORMATION FOR SEQ ID NO: 66:	
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 1011 base pairs	
50	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
50	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 66:	
55	CGGAAGAAAG CAGCCATCCA GACATTTCAG AACACGTACC AGGTGTTAGC TGTGACCTTC	60
	AATGACACAA GTGATCAGAT TATTTCTGGT GGAATAGACA ATGATATCAA GGTCTGGGAC	120

TGCGCCAGAA CAAGCTAACC TACACCATGA GAGGCCATGC AGATTCAGTG ACTGGCCTGA

	GTTTAAGTTC TGAAGGCTCT TATCTTTTGT CCAATGCAAT GGACAATACA GTTCGTGTCT	240
	GGGATGTCCG GCCATTTGCC CCCAAAGAGA GATGTGTAAA GATATTTCAA GGAAATGTGC	300
5	ACAACTTTGA AAAGAACCTT CTGAGATGTT CTTGGTCACC TGATGGAAGC AAAATAGCAG	360
	CTGGCTCAGC CGACAGGTTT GTTTATGTGT GGGATACCAC AAGCAGGAGA ATATTGTATA	420
10	AGCTGCCCGG CCATGCTGGC TCCATCAATG AAGTGGCTTT CCACCCTGAT GAGCCCATCA	480
- 0	TTATCTCAGC ATCGAGTGAC AAGAGACTGT ATATGGGAGA GATTCAGTGA AGATATGGAC	540
	TGGAAGACTC CAAGGCCGCT TGTCTTTGAG ACCTCAGACT GCATAAGTGA TGCCAAATGT	600
15	TGGATGTCCA GGYTAGCACC CTCCCTTCAG ATGACCATTG CTAGCAAGAA ACAGGAGGCG	660
	GTGGCCATAT TCCAAAAACC ACTTCTGTCC CATTTCACCA GGATGACTAA GGCAAGCTCC	720
20	CTGTGGCCTC TAAAAACCAC CTGCCAGATT TCAGGGACTG TTTTTTTTTT	780
	TITTCCTGTT TTCTAATGCA GGCCCAATGT GACAAATTTG TTGGTTGGGA TTTTTTTTTT	840
	TIPITGTAAC TGGCTTGTAT GATATTTTCT TTCTGTATTT CTCTATATCA TTTTGTATTA	900
25	AAAGCCAAAT AGATGCCTTT TTACAAGARM AAAAAAAAAA AAAAAAAAA NNAAAAAAAA	960
	CTGGGAGGGG GGGCCCGGTA CCCAAATCGC CGGATATGAT CGTAAACAAT C	1011
30		
	(2) INFORMATION FOR SEQ ID NO: 67:	
35	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1193 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 67:	
	GGCCGGGCGG TGCGCACTGC GGGCGCATCC CTGCCCCGGC GCCGTCCGTG CCCGCGGGAC	60
45	CTGACAGCCG GGTCAGAGGG CGAACTGTGC TCAGGCCCGG GCTGGACGCA GAGCCAGAGC	120
.5	TGTCCCCAGA GGAGCAGAGG GTCCTGGAAA GGAAGCTGAA AAAGGAACGG AAGAAAGAG	180
	AGAGGCAGCG TCTGCGGGAG GCAGGCCTTG TGGCCCAGCA CCCGCCTGCC AGGCGCTCGG	240
50	GGGCCGAACT GGCCTGGGAC TACCTCTGCA GATGGGCCCA AAAGCACAAG AACTGGAGGT	300
	TTCAGAAGAC GAGGCAGACG TGGCTCCTGC TGCACATGTA TGACAGTGAC AAGGTTCCCG	360
55	ATGAGCACTT CTCCACCCTG CTGGCCTACC TGGAGGGGCT GCAGGGCCGG GCCCGAGAGC	420
- -	TGACGGTGCA GAAGGCGGAA GCCTGATGCG GGAGCTGGAT GAGGAGGGCT CTGATCCCCC	480
	CCTGCCGGGG AGGGCCCAGC GCATCCGACA GNTGCTGCAG CTGCTCTCCT AGTGGGTTCA	540
60	GCGCGGGGCG GGGCCGCTGC CCAGTGCAGG GCTGCCTCAG ACCACACAGG GTGCAGCTCC	600

	TCCGGCGGTG GGGGCCGGGT TCACCAGCAG GGCAGCGGCT GAGCAAGGGC TTTCAGCTCC	660
_	TCCGGTGGTG GGGGCCGGGA TCACCAGCAC CAGAGCCTCG CAAGGGCCCC TTCCCTCCTC	720
5	CAGACCCTCC TTGGCCGGTG ACGCTGTGAC AGTGATGGCA GGTTCAGTGC CTTCAGCGCA	780
	GAGCGTGGAT GCTCTGGAAT CACCCGGACC CCTGGCCTTG GAGGGACCCT CCAGCCCCAG	840
10	GAATCTGCTT TGGAGGGAAA TGTCTATTTT TCTACCGGGA ATATTTTAGA GATTGGGGCA	900
	TGCTGGCTCC TCCCGCCAGC TGCAAACCTG CACCTTCCGC CTGATTCCCG ATCCCCCTGC	960
. ~	GTGGGCCGCA TTCCTGGTCC CCTGCCTGCG TCCATCGAGG GGCCTGGCTG TGGCCTGTTT	1020
15	TCCTTTGACC CCACACAGCG TCATTGCGGG TCATGGGGAG CCCCTGGTGG GAGCTTGTGG	1080
	AGTCGGATCA CGTACCTGTG CAGAAACCGC CTCTGTGGCT GCATTTGAAA TAAAACCCGA	1140
20	CCCAGCAGCA AAAAAAAAA AAAAAANCNC NAGGGGGGGC CCGGNACCCA ATT	1193
25	(2) INFORMATION FOR SEQ ID NO: 68:	
20	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 560 base pairs	
30	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 68:	
35	GAATTCGGCA CGAGTTGGCA CATGATGCAA AATGCATTTC TCAGAGTAGA TTGCAGTCAA	60
	AAATGTTGGA AACTACTAAG CATGTGCARA TAGCATGCAT GCTGCTGCTG ACCTGCCAGA	120
40	TATTTCTCCC TTCCTCCCTT TCTCCCTCAT TTATTCATTC	180
40	CATTAAAAAA ATTATATGTA TGTTTTGTGC AAAGCACCCT ACTCAAGGCT GCGGGGTACA	240
	AAAGTATATC AGAAGCCTTG GGCTTTGACM WACTTCTCTG TAGTAGTGCT AGATTTGTGT	300
45	GGATCTGCCA CACTTACTCC AGGCCTCTTG TGACCTGTGC TTTGCATTAA TCTCTTAGGC	360
	TAAGCCACAT ACCTITICAT TATACAATCT TIGCTGATGC TAAGGACAGA TICCAAAGTG	420
50	CCCTCCTTAT AATTTTTGTA TTTAATGCAA AGTGTAATCA AGAATAGGCC ATTGTTAGGT	480
50	CAATTGCTTT TCTGTATTTA TCTTTTCAAA CAATAAATAA TCAGTGGGAT GAAAAAGGGC	540
	CGGAAAAAA AAAAAAAAA	560

(2) INFORMATION FOR SEQ ID NO: 69:

60 (i) SEQUENCE CHARACTERISTICS:

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(A) LENGTH: 1657 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 69: CGGACNGAGC CGCCGCCGGG CACTTCCTGT GGAGGCCGCA GCGGGTGCGG GCGCCGACGG 60 10 GCGAGAGCCA GCGAGCGAGC GAGCGAGCCG AGCCGAGCCT CCCGCCGTCG CCATGGGCCA 120 GAACGACCTG ATGGGCACGG CCGAGGACTT CGCCGACCAG TTCCTCCGTG TCACAAAGCA 180 GTACCTGCCC CACGTGGCGC GCCTCTGTCT GATCAGCACC TTCCTGGAGG ACGGCATCCG 240 15 TATGTGGTTC CAGTGGAGCG AGCAGCGCGA CTACATCGAC ACCACCTGGA ACTGCGGCTA 300 CCTGCTGGCC TCGTCCTTCG TCTTCCTCAA CTTGCTGGGA CANTGACTGG CTGCGTCCTG 360 20 GTGTTGAGCA GGAACTTCGT GCAGTACGCC TGCTTCGGGC TCTTTGGAAT CATAGCTCTG 420 CAGACGATTG CCTACAGCAT TTTATGGGAC TTGAAGTTTT TGATGAGGAA CCTGGCCCTG 480 GGAGGAGGCC TGTTGCTGCT CCTAGCAGAA TCCCGTTCTG AAGGGAAGAG CATGTTTGCG 540 25 GGCGTCCCCA CCATGCGTGA GAGCTCCCCC AAACAGTACA TGCAGCTCGG AGGCAGGGTC 600 TTGCTGGTTC TGATGTTCAT GACCCTCCTT CACTTTGACG CCAGCTTCTT TTCTATTGTC 660 30 CAGAACATCG TGGGGCACAG CTCTGATGAT TTTAGTGGCC ATTGGTTTTA AAACCAAGCT 720 GGCTGCTTIG ACTCTTGTIG TGTGGCTCTT TGCCATCAAC GTATATTTCA ACGCCTTCTG 780 GACCATTCCA GTCTACAAGC CCATGCATGA CTTCCTGAAA TACGACTTCT TCCAGACCAT 840 35 GTCGGTGATT GGGGGCTTGC TCCTGGTGGT GGCCCTGGGC CCTGGGGGTG TCTCCATGGA 900 TGAGAAGAAG AAGGAGTGGT AACAGTCACA GATCCCTACC TGCCTGGCTA AGACCCGTGG 960 40 CCGTCAAGGA CTGGTTCGGG GTGGATTCAA CAAAACTGCC AGCTTTTATG TATCCTCTTC 1020 CCTTCCCCTC CCTTGGTAAA GGCACAGATG TTTTGAGAAC TTTATTTGCA GAGACACCTG 1080 AGAATCAATG GCTTCAGGAC ATGGGTTCTC TTCTCCTGTG ATCATTCAAG TGCTCACTGC 1140 45 ATGAAGACTG GCTTGTCTCA GTGTTTCAAC CTCACCAGGG CTGTCTCTTG GTCCACACCT 1200 CGCTCCCTGT TAGTGCCGTA TGACAGCCCC CATCAAATGA CCTTGGCCAA GTCACGGTTT 1260 50 CTCTGTGGTC AAGGTTGGTT GGCTGATTGG TGGAAAGTAG GGTGGACCAA AGGAGGCCAC 1320 GTGAGCAGTC AGCACCAGTT CTGCACCAGC AGCGCCTCCG TCCTAGTGGG TGTTCCTGTT 1380 1440 55 GATAAGTGGG ATCTACCAAT TGATTCTGGC AAAACAATTT CTAAGATTTT TTTGCTTTAT 1500 GTGGGAAACA GATCTAAATC TCATTTTATG CTGTATTTTA TATCTTAGTT GTGTTTGAAA 1560 60 1620

	AAAAAAACTC GRGGGGGGC CCGGTACCCA AATCGCC	1657
5		
	(2) INFORMATION FOR SEQ ID NO: 70:	
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 711 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 70:	
	GGCACGAGCG AAGACCCTGT TCGGACCCTG CCCCGATTCC AGACTCAGGT AGATCGTCGG	60
20	CATACCCTCT ACCGTGGACA CCAGGCAGCC CTGGGGCTGA TGGAGAGAGA TCAGGTATCC	120
20	CCCAGGGAGT AGGGGCTACC TTGAGGGGAT GATAGACCTC CCCCACTCCC AGTGKKACTC	180
	TGGAAATATG AAGGAACTAG GGAGTGGAAG AGATTTCAGA GCTGGGGAGA GGAGTTCCTC	240
25	CCTTCAAAGC CAGCAACTGC CTTTGGGGAA TGTCGGGGGG TCTCTCCTTT CTCCTGCTTG	300
	TTTRAGGTGG TACACAGTCC CCCCTTCAMC TGGSGGGAAG CTGTNCCGGA CARACTCATC	360
30	TCAGCTTTCC CTTGGGGCAG GATCGGGGGC AGCAGCTCCA GCAGAAACAG CAGGATCTGG	420
30	AGCAGGAAGG CCTCGAGGCC ACACAGGGGC TGCTGGCCGG CGAGTGGGCC CCACCCCTCT	480
	GGRAGCTGGG CAGCCTCTTC CAGGCCTTCG TGAAGAGGGA GAGCCAGGCT TATGCGTAAG	540
35	CTTCATAGCT TCTGCTGGCC TGGGGTGGAC CCAGGACCCC TGGGGCCTGG GTGCCCTGAG	600
	TGGTGGTAAA GTGGAGCAAT CCCTTCACGC TCCTTGGCCA TGTTCTGAGC GGCCAGCTTG	660
40	GCCTTTGCCT TAATAAATGT GCTTTATTTT CAAAAAAAAA AAAAAAAAAC T	711
45	(2) INFORMATION FOR SEQ ID NO: 71: (i) SEQUENCE CHARACTERISTICS:	
50	(A) LENGTH: 935 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 71:	
	GCCACAGGGT GAAAGCCAGC TAAACCCCAA GTGGAGAAGT GAAAGACATG GTTGTTCCCA	60
55	TAAGTTTATT GCTCACATTA TGAAAGAAGC CATAGTCATG AGTGAACCAC TCCCTAGGTT	120
	TAAGTITATT GCTCACATTA TGAAAGAAGC CATAGTCATG AGTGAACCAC TCCCTAGGTT GATAAGGAAA CCAACACGGA AGATCTCTTT CTGGAAGAAG CAGCCAGCCT CGTGAAGGAG	180
60		240
OU	CACCCARGO ACCARGO NAVALCACCI III O I COMI O I COMI I CONTROLLI	'

	CAGACATTCT CGCCTGGAGC ACGAAGCCAG TATGTTTGCA GACTTTATCG TAGTGACAGC	300
5	GACAGTTCAA CGCTGCCCCG GAAGTCCCCC TTTGTCCGAA ATACTTTGGA AAGACGAACC	360
	CTTCGCTATA AGCAGTCATG CAGGTCTTCC CTGGCTGAGC TCATGGCCCG CACCTCCCTG	420
	GACTTGGAGC TGGATCTCCA GGCGTCGAGA ACACGGCAGA GGCAGCTGAA TGAGGAGCTC	480
10	TGCGCCCTCC GTGAGCTGCG GCAGCGGTTN GGAGGACGCC CAGCTCCGTG GCCAGACTGA	540
	CCTCCCACCC TGGGTGCTTC GGGACGAGCG GCTCCGTGGC CTGCTGCGGG AGCCGAGCGG	600
15	CAGACAAGAC AGACCAAACT TGACTACCGT CATGAGCAGG CGGCTGAGAA GATGCTGAAG	660
	AAGGCCTCCA AGGAGATCTA CCAGCTGCGT GGCAGAGCCA CAAAGAGCCC ATCCAAGTGC	720
	AGACCTTTAG GGAGAAGATA GCATTCTTCA CAAGGCCAAG GATCAACATA CCTCCTCTCC	780
20	CAGCCGACGA CGTCTGATGG AGTGCATTGT GCACATGAAG TATTTATCCA CCTGTTTTAT	840
	TTTCATGAAG TTCTTAGACT AGCTGAATTT GTCTTTAAAA TATTTGTGCA AAGCTATTAA	900
25	TATACACATT TTGTAAAAAA AAAAAAAAAA AAACT	935
•	(2) INFORMATION FOR SEQ ID NO: 72:	
30	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 504 base pairs(B) TYPE: nucleic acid	
35	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 72:	
40	GCAGGGGCGA GGGGYTGGGG ACCGCGGGGC GGACGGGAGC GAGTATGTCC GCTCTGACTC	60
40	GGCTGGCGTC TTTCGCTCGC GTTGGAGGCC GCCTTTTCAG AAGCGGCTGC GCACGGACTG	120
	CTGGAGATGG TGGAGTCCGT CATGCCGGTG GTGGTGTGCA CATTGAGCCC CGGTATAGAC	180
45	AGTTCCCCCA GCTGACCAGA TCCCAGGTGT TCCAGAGCGA GTTCTTCAGC GGACTCATGT	240
	GGTTCTGGAT TCTCTGGCGC TTTTGGCATG ACTCAGAAGA GGTGCTGGGT CACTTTCCGT	300
- 0	ATCCTGATCC TTCCCAGTGG ACAGATGAAG AATTAGGTAT CCCTCCTGAT GATGAAGACT	360
50	GAAGGTGTAG ACTCAGCCTC ACTCTGTACA AGAGCCAGGT GAGAATTTCA AGGATTATCG	420
	ACTICATATI GCACATTAAA GITACAAATI AAAGIGGCII GGICAAGAAI GARAAAAAA	480
55	AAAAAAAATT GGGGGGGGC CCCN	504
		JU-1

(2) INFORMATION FOR SEQ ID NO: 73:

5	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 620 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 73:	
10	GAATTCGGCA CGAGGAGGAG GGGAGGCGGG GTAAGTTTGG TGGGAAACTC TGTAATTTCC	60
	WITTITACTT TCACAGCAAT AGTGCAGAAT CCAGAATGGA TGTCCTCTTT GTAGCCATCT	120
1.5	TTGCTGTGCC ACTTATCCTG GGACAAGAAT ATGAGGATGA AGAAAGACTG GGAGAGGATG	180
15	AATATTATCA GGTGGTCTAT TATTATACAG TCACCCCCAG TTATGATGAC TTTAGTGCAG	240
	ATTTCACCAT TGATTACTCC ATATTTGAGT CAGAGGACAG GCTGAACAGG TTGGATAAGG	300
20	ACATAACAGA AGCAATAGAG ACTACCATTA GTCTTGAAAC AGCACGTGCA GACCATCCGA	360
	AGCCTGTAAC TGTGAAACCA GTAACAACGG AACCTCAGAG TCCAGATCTG AACGATGCCG	420
25	TGTCCAGTTT GCGAAGTCCT ATTCCCCTCC TCCTGTCGTG TGCCTTTGTT CAGGTGGGGA	480
25	TGTATTTCAT GTAGAAGGTG GAAGAAGGCT GCTATGACTC TTTGGATGGG AGTCTGGCAA	540
	GAGGAAATTG GAAGATAAAA TAAATAATAA GTGAAATAAA AAAAAAAA	600
30	GGGGGGCCC GGTACCCAAT	620
35 40	(2) INFORMATION FOR SEQ ID NO: 74: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 581 base pairs (B) TYPE: nucleic aci (C) STRANDEDNESS: double (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 74:	
45	ACAAGGTGTG TGTAAAGTTT ATGTTTGTAA ACTGAATTCT ATCTTAAATC CAAAAAGAAC	60
43	TCGGGAGTAA TTCATTTTTG TAGCATAAAG ATCCCTAAGT TTTATTTTGA AATATCTGAT	120
	TTTTACACGT TAAAAAATAA CAGGGCATCG AGAGGATTCC TAGGTGACAT CCAGACTCCT	
50	TTAGCTTTGT GTGTGTGGCA CCGGTTAGTC TGCTTCTCTC TCCTTTCTTG CACTGCTTCA	240
	CACAGCCATG CCCTGCCAGC CCGGGCAGGT GCCTTCCTGT CAATGTACAT TTGGGCTTCT	
55		
55		
	GGAGTACTTC CATGGGTGTG CCTCCCCCAG CCAAGCCATA ATAGGTGGTT TCCCCTTCGC	480
60	TTCTGTAGCC CTTGCAGACA TCCTCTGTTT ACAGTAGGTG TTGACTTACT TCCCCTCTCC	#00

	CCGSTAAAGC	CATAAACTCC	TTAAGGACAG	GTAGCATTCT	TAGTATCTTC	GTTCTTCTCA	540
	ATGACCAGTA	GACCATTAAA	CATGTAGCAA	ACAAATGTGA	A		581
5							
	(2) INTEGRAL	ATION FOR SI	FO TO NO. 75	· .			
10			_				
15	(1)	(B) TYP (C) STR	GTH: 1843 b E: nucleic ANDEDNESS: OLOGY: line	ase pairs acid double			
13	(xi) SEQUENCE	DESCRIPTION	: SEQ ID NO	: 75:		
	AAACCCAACN	CCCTCCGGTC	CCCNAAAGAA	AGCCCAGCCC	AAATCCCAAG	CCGGCAGTGA	60
20	GCCCGCGAAC	AAGGCCCTCA	AGACGCCCAG	NCGAACAAGC	AGCCCCCAGG	AGGCCCCGCA	120
	AGAGAACTCC	CTGGCGGCCC	AAGCGGCAG	CTTCTGTGCG	GCAGAACTCA	GCCACCGAGA	180
25	GCGCAGACAG	CATCGAGATT	TATGTCCCGG	AGNCCCAGAC	CAGGCTCTGA	GACCATGCAG	240
23	GAGGAAAGAA	ACGATTTTAA	ATCATTAAAA	ACACAAAAAC	TAAGTGCGAA	CGGAACAGAG	300
	TTTTCTCAAC	CTTTGCTATG	GTTATTCTGT	CTAGAGACCC	TGAGCCAACT	TTCAAATTGA	360
30	CGCATACAAG	GGCTCACAAT	TTGGCTTTTT	TGGGTCCCTC	CCAGCTTTAG	GTTATGAAGA	420
	TTTTACTCAC	AAAAAAATC	AACAAAAATC	ACGAAACTAG	AAAACTTTTT	TTTTCCTCTT	480
35	GCTGGCCGTG	GTGGACTAGA	TAGATGGACG	TCGGCAACTC	CCGGCCCAGC	CTCCATACTG	540
	CGGTCTTTTT	ACTCGTTCTA	TCTGATGAGA	ACTONINGTA	GCTTGTTTAC	AAGATGACGA	600
	CAGTCCAAGG	GCAGCCTTGG	GCACCTGCCA	IC -	TTCCCCAGCT	ATCCCCCCTC	660
40	TGACCTTGAT	TTTCATTCTT	ATGTTTTTCT	C.PT	CAGAGCTCAC	ACAGTGGTCA	720
	CCATTGTGGC	AAGCGGCTTT	CTGGGTCTCA	GCCTCTCTG	CGGTTGAGGG	CCCAGAGGAC	780
45	AGAGAGATGG	ACATGCGTCC	CCTCCCTCCC	CCCGCCAAGT	GCTCACACAC	AACCTCACGC	840
	GCACACACAC	ACACGCAGAT	GGAGGCGCCT	CACTGGGAGG	TGCCCCGCCA	GCCCTGGGCA	900
	GTGTCAGGCA	GGACTCACTC	ACCGCTGAGC	AGATGAGAGA	AGTTTTAGTC	TTGGCGGGTG	960
50	GAAATGAGAC	GAAGCCACAG	TTATCACACT	CCAGACTCCT	GCCCTTTTAT	TTTCTCCAGC	1020
	CCCTTCTTCC	TTCAGCAAAA	TCTAGGACTC	CCGAGTGGCT	TCCAGGGGC	CGTCAGTCCT	1080
55	CAGCCGCGCC	TGTGTCCGGT	GCCCGAGGGG	ceececec	TGTCTGTATG	TATGTGTACA	1140
	TATGCACATA	GACCTTAGAG	TGTATAGTTA	ACAAACGCCC	ATCTGCTCAC	CCATGCCCAC	1200
	CCAGCGCCGC	CGCCGCTGGC	TCTCGGGGCA	CCTGGCAGGA	GCCGGTGTG	TGAATAGCAT	1260
60	ATATTTTAC	ATGTACTATA	TCTAGGTGTG	TGTACAAGTG	TGTGTAAAAA	TATATACCTT	1320

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	GTGTGTAAGC	AGCCCTTTTT	TTTTTTGGTC	TCCACCCCCC	TCCCCCCGCC	CCGCACTCCT	1380
5	AAGGGCCCAT	CTGCCCAGCC	TCTGAGTTTT	CTGTTCTATT	TTTTTTTAA	CCCCAATTAT	1440
3	CCTTCTCTCT	CTCCTGCCCC	CGCATCCCAC	TCCCAGGGTG	TCACGAGCCC	TGAGCTGCAA	1500
	TGGCCCGGGC	CTGCAGGGCG	GGGTAGGGGA	GGGCARGGCT	SAGCCCCGAA	GCCAGCTCAG	1560
10	TACCTGAGGG	GCTGCTCTAT	GCTGTGTATG	CGCCTCTCTG	GCATCCGAGA	CATCCTCTTG	1620
	GTGGCGCTTG	CTNGCAGGGG	ACCCCCCCC	CGTCCCCAGG	TGAACCAAGG	GTCTGCTCCG	1680
15	GGGCCCATTT	CCAGCTTGGC	CGCCGTCTGT	GACCTTGGGC	AAGTCACTTG	ACCTCTGTGT	1740
13	GCCTCAACTT	CCTCCTCTGT	AAAACGGGGA	CAGTCCCTGC	CCCTCCCTAC	CTCACAGGCA	1800
	TGTTGTGAGA	ATAAATGAGG	TAACGTGTAA	АААААААА	AAT		1843

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(2) INFORMATION FOR SEQ ID NO: 76:

25 (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1441 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

30

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 76:

	TCGACCCACG CGTCCGGCTC CCCGAGCCCT GCCAACCATG GTGAACTTGG GTCTGTCCCG	60
35	GGTGGACGAC GCCGTGGCTG CCAAGCACCC GGGACTCGGG GAGTATGCCG CATGCCAGTC	120
	ACACGCCTTC ATGAAGGGCG TTTTCACCTT CGTCACAGGC ACCGGCATGG CCTTTGGCTT	180
40	GCAGATGTTC ATTCAGAGGA AGTTTCCATA CCCTTTGCAG TGGAGCCTCC TAGTGGCCGT	240
40	GGTTGCAGGC TCTGTGGTCA GCTACGGGGT GACGAGAGTG GAGTCGGAGA AATGCAACAA	300
	CCTCTGGCTC TTCCTGGAGA CCGGGCAGCT CCCCAAAGAC AGGAGCACAG ATCAGAGAAG	360
45	CTAGGAGAGC TCCAGCAGGG GCACAGAGGA TTGGGGGCAG GAGGAGTCTG GAACACAGCC	420
	TTCATGCCCC CTGACCCCAG GCCGACCCTC CCCACACCCT AGGGTACCCC AGTCGTATCC	480
50	TCTGTCCGCA TGTKTGGCCA GGCCTGACAA ACACCTGCAG ATGGCTGCTG CCCCAACCTG	540
50	GGACCTGCCC AGRAGGTTGG AGCAGAAAGG GCTCTCCCTG GGGTGGTGTT TCTCCTCTAG	600
	GGTATTGGGA TGCATGTTCT GCACTGCCAG CAGAGAGGGT GTGTCTGGGG GCCACCACCT	660
55	ATGGGACACG GGGTCGAAGG GGCCTGTACA CTCTGTCATT TCCTTTCTAG CCCCTGCATC	720
	TCCAACAAGT CCAAGGTGAC AGCTGGTGCT AGGGGCGTGG GGTTAATAAA TGGCTTATCC	780
60	TTCTCTCCAC CCAAGTTTCC ACCTGACCAG GTGAAAAACA AATCAGAAGG GTAAGATGAT	840

	GACAGGTCAC ATGAAACCTT TATTACCCTA CAGTTGATAT ATGAGGATCA CATGCAAGTT	900
	ACATACTGAG GATGTACAGG GAAGTTCCCA GCGCTGAACC CCAGAATTAG ACGTTCGCAT	960
5	CAGCCCCGTA GGCCACGTGG ACACCACCAC AGCCTCTCTG TATGGGGGTC TGCCTCTGTA	1020
	GCACTTGGCA TGTAGGGGCA GAGCAAAAGG GGCCANGCTG GCCAGAGCCT GGCTGCTGGG	1080
10	NAGARGAGGG ACTIGIGGGS CACGCCACNI GCCTATCATI CCCCAYICAI CTATIAGCCA	1140
	AAGTCACTCC CCAGAGGCAG AGCTAGCCCG TTGTAGCCGT GTCTGTGTGG AGGGAAAGCT	1200
	TCTGAGTGGG CAAGCCTACA CACAGCCCCG AGCCCCAAGA GGAGGAAGAG GTGGAGACCA	1260
15	GACGGAACCT CCACAAGTCC ATCATGGTTA CAGCTGGCTT CCCCGCAGCA CCGAAGACCC	1320
	ACAGCATNGG CCCTGCTGCC CCCGACCCAG CTCAGCTGCC ANGCCTCACC TTGCCAGGAA	1380
20	TTGAAAGAAA GTTATTGAGT ACTAATTGGC CTCAGAGTNA CAGGAAGCTC AAGTTAAAGT	1440
	G	1441
25	(2) INFORMATION FOR SEQ ID NO: 77:	
	(i) SEQUENCE CHARACTERISTICS:	
30	(A) LENGTH: 910 base pairs (B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 77:	
35	GGCAGAGCTG GCCTTCGACT CGCTATGTCC ACTAACAATA TGTCGGACCC ACGGAGGCCG	
	AACAAAGTGC TGAGGTGAGG ACCCCAGCGT CGTGGGCACG GGTTCGGGTT GTGGGTGTGG	60
40		120
	ATCGGGGCCC TGGGAAGCGC CTGTCTATCC CGGGGGCAGG ACCTGAGCGC CCCTGACCCT	180
	CGAGCCTGTC GCAGGTACAA GCCCCCGCCG AGCGAATGTA ACCCGGCCTT GGACGACCCG	240
45	ACGCCGGACT ACATGAACCT GCTGGGCATG ATCTTCAGCA TGTGCGGCCT CATGCTTAAG	300
	CTGAAGTGGT GTGCTTGGGT CGCTGTCTAC TGCTCCTTCA TCAGCTTTGC CAACTCTCGG	360
50	AGCTCGGAGG ACACGAAGCA AATGATGAGT AGCTTCATGT GAGACTTGCC CTACAGAACA	420
50	AGTGACTCTT GAGTAAGGGG TGGGGGGACC CCAGCCTGGC CATCCTAGAC TGACACCTCT	480
	CTCCTGTCTT CATGCTGTCC ATCTCTGCCG TGGTGATGTC CTATCTGCAG AATCCTCAGC	540
55	CCATGACGCC CCCATGGTGA TACCAGCCTA GAAGGGTCAC ATTTTGGACC CTGTCTATCC	600
	ACTACCCORC CCCMMMCCCCM COMMAND	
	ACTAGGCCTG GGCTTTGGCT GCTAAACCTG CTGCCTTCAG CTGCCATCCT GGACTTCCCT	660

CACCCTAGGC TTACCCCTCC TGCCTCCCTT CCCCTGCCTG CTGCTGGGGG AGATGCTGTC

1140

1200

	CATGITITCTA GGGGTATTCA TITGCTTTCT CGTTGAAACC TGTTGTTAAT AAAGTTTTTC	840
_	ACTCTGAAAA AAAAAAAAA AAAAAAAAAC TYGRGGGGGG GCCCGGAACC CAATTCSCCG	900
5	GATAGTGAGT	910
10	(2) INFORMATION FOR SEQ ID NO: 78:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 2776 base pairs	
15	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
	(D) TOPOLOGY: linear	
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 78:	
20	TCGACCCACG CGTCCGGGCG GGCAGTGATG GCGGCTGGTG ATGGGGACGT GAAGCTAGGC	60
	ACCCTGGGGA GTGGCAGCGA GAGCAGCAAC GACGGCGGCA GCGAGAGTCC AGGCGACGCG	120
25	GGAGCGGCAG CGRAAGGGG AGGCTGGGCG GCGGCGGCGT TGGCGCTTCT GACGGGGGC	180
	GGGGAAATGC TGCTGAACGT GGCGCTGGTG GCTCTGGTGC TGCTGGGGGC CTACCGGCTG	240
20	TGGGTGCGCT GGGGGGGCG GGGTCTGGGG GCCGGGGCGG GGGCGGCGA GGAGAGCCCC	300
30	GCCACCTCTC TGCCTCGCAT GAAGAAGCGG GACTTCAGCT TGGAGCAGCT GCGCCAGTAC	360
	GACGGCTCCC GCAACCCGCG CATCCTGCTC GCGGTCAATG GGAAAGTCTT CGACGTGACC	420
35	AAAGGCAGCA AGTTCTACGG CCCGGCGGGT CCATATGGAA TATTTGCTGG TAGGGATGCC	480
	TCCAGAGGAC TGGCCACATT TTGCCTAGAT AAAGATGCAC TTAGAGATGA ATATGATGAT	540
40	CTCTCAGATT TGAATGCAGT ACAAATGGAG AGTGTTCGAG AATGGGAAAT GCAGTTTAAA	600
40	GAAAAATATG ATTATGTAGG CAGACTCCTA AAACCAGGAG AAGAACCATC AGAATATACA	660
	GATGAAGAAG ATACCAAGGA TCACAATAAA CAGGATTGAA CTTTGTAAAC AACCAAAGTC	720
45	AGGGGCCTTC AGAACTGCAA TTCTTACTCC CTTTCACAGA CTGTCCGGAG TCTTTGGGTT	780
	TGATTCACCT GCTGCGAAAA ACATTCAACA AATTGTGTAC AAGATAAATT AATCTCACTA	840
50	TGAAGATTTG AATAACTAGA CATTATTTAT GCTGCCAAAC TCATTTGTTG CAGTTGTTTG	900
50	TAATGTCTAG TGGGGCTTCA TCATCCTGAA AAGAAGGAGA CAGGGATTTT TTTAAAGAGC	960
	AAGAAAGTCA CAATATTACT TCTTTCCTTC CTTTTTTCCT TCTTTCCTTT CTTCTT	1020

TTTCTTTCTT TTTAAAATAT ATTGAAGACA ACCAGATATG TATTTGCTAC TCAAGTGTAC

AGATCTCCTC AAGAAACATC AAGGGACTCC TGTGTCACAT ACTGTGTTTT TATTTTAACA

TGGGTGAGGG AGGCGACCTG ATCAGGGGAG GTGGGGGTAC ACATCAATTT GAGTTGTTCA

55

	GGCTACTGAA	ACATTAAAAT	GTGAATTCCC	AAACTTTTCT	TTTTGGCTTT	GTCAGGGAAA	1260
	AGAAAAATAT	CTTTATAAAG	AAATCTTTGG	AAATTAGGAG	AAGGAATTTC	AGGTGGGTTT	1320
5	AAGTCAGAGC	TAGTTCCCCA	ACAGAAAGAT	CATTTGAAAC	CAGTTTTTAT	CCCTTCTCTT	1380
	TCCTTCCCTT	TCCCTAAATC	AAATCAATAT	TAATTGTGCC	TTATTTCACT	TAACATAGAC	1440
10	TTGAATTATT	TTTAGGGAAA	GCCCCTATAA	TGAATTCAGA	AATCACTACA	AGCAGCATTA	1500
	AGACTGAAGT	TGGAATATTC	TGTTGACCAT	AAAACCTTGA	TATCATTCTG	TGTATATAGA	1560
	ATGTAAAAGG	AATATTACAG	TGTTAACTGC	CATATATGTA	ATATACACAA	ACTCAATTAG	1620
15	CATTGTAATG	GCCAAATGCA	TTCCCCCATG	CTTTTCTGTT	TTCAAAAAA	TTGAAAAACA	1680
	AATCAACTCT	TATCCCCAAC	AGCTGCCTAA	TTTTAGGAGT	CTGACCCTCC	ACATCTCACT	1740
20	GGTGTGGGTG	CATGGGGCTG	TGGAGTGGGT	GTCAGTATGG	ATGTGTCTGA	ATGTGTGAGG	1800
	CCTTGGAAGG	GACTCTTTCT	GCAGATACTG	TAAATACAAG	TACCATTTTA	ATAAAGCATG	1860
	TACAATAAAC	CAAAATAAGC	TTGAGTTGGA	CTTTATATAC	AGAACTGTAA	GCCAGTGCAT	1920
25	TATGATACAG	TTGTAAGATT	GTGCATTTGA	TTCAAGATAA.	GGAAAAATCT	TGGAAATGAA	1980
	AAGCAGGCAC	KGGTTAACCA	AGTTGTACAC	ATTGTACCAC	ATTCAGCATA	ACTTTAGGAA	2040
30	GAAATTCCAC	TTTGTGAACA	TTCTCCAGAA	ATCCAAGATT	ATTCAGGTAA	GAATTGGTAT	2100
	ATTAAATGTA	CATCTTTTTA	CTTTCTATTT	TGATGCCAAC	TGATTATACT	AGACAATTAG	2160
	CACTCCAGGT	GGTTATTGAA	CACAAAACAG	TAAAAGAATA	TTGCACTGAT	AGATACTAAA	2220
35	TTATTATTTT	ATTAGGTTGA	AAAAGCCCTT	ACTAAAAGCC	CCTCATATAT	CAATTACTTT	2280
	ATTTCATTAT	GACTACTTAG	GTTCCGGGCT	GGGGACAAGT	TCACTTAAAA	AGGCAATGTT	2340
40	ATTTAACAGG	TCACCAGTTA	AGACTTCTGC	TTTGTAGATA	CATGCAGAAG	CCATCAAACA	2400
	AGGGGGRGCT	TTTAACTGCA	ACAATAAGCT	AAAGTATGTA	AAATACTACA	TTCTATTCAG	2460
	TCTTGGAGTG	TTTTGTAGAA	AGTTATCTTC	AGCCAAATCT	TTGCTGAAGA	CTGGTTGTGG	2520
45	AGTGTTGGTA	AATGCTTTGT	GTTTTTATGT	AAAATATTTT	СТАААСАААА	AATGTTAAAA	2580
	GTACATGTCC	TCTGTAGTAA	ACTGATATCT	ATATATATGA	ATCATTCAAG	CCTAAAGTCT	2640
50	AGTAATAAAC	TGTACTTGTG	AATAGAGAAA	СССТАААТАТ	TCATGCAGWA	AAAATTATGC	2700
	GGTCTGTTAA	GAAAAATGAG	TAATTTGTGT	TTTGGACTTG	AAATAAACAG	TGTTCTGTAG	2760
	ATAATTCCTC	AACTTC					2776
55							

(2) INFORMATION FOR SEQ ID NO: 79:

60 (i) SEQUENCE CHARACTERISTICS:

WO 98/42738 PCT/US98/05311

229

(A) LENGTH: 1525 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear

5

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 79: CCGCTGCTGA TAACTATGGC ATCCCCCGGG CCTGCAGGAA TTCGGCACGG AGCTACGGCG 60 CCGCCTGGCT CCTGCTGNCA CCTGCAGGCT CGTCGCGGGT GGAGCCCACC CAAGACATCA 120 10 GCATCAGCGA CCAGCTGGGG GGCCAGGACG TGCCCGTGTT CCGGAACCTG TCCCTGCTGG 180 TGGTGGGTGT CGGCGCCGTG TTCTCACTGC TATTCCACCT GGGCACCCGG GAGAGGCGCC 240 15 GGCCGCATGC GGASGAGCCA GGCGAGCACA CCCCCCTGTT GGCCCCTGCC ACGGCCCAGC 300 CCCTGCTGCT CTGGAAGCAC TGGCTCCGGG AGCSGGCTTT CTACCAGGTG GGCATACTGT 360 ACATGACCAC CAGGCTCATC GTGAACCTGT CCCAGACCTA CATGGCCATG TACCTCACCT 420 20 ACTCGCTCCA CCTGCCCAAG AAGTTCATCG CGACCATTCC CCTGGTGATG TACCTCAGCG 480 GCTTCTTGTC CTCCTTCCTC ATGAAGCCCA TCAACAAGTG CATTGGGAGG AACATGACCT 540 25 ACTICICAGG CCICCIGGIG ATCCIGGCCI TIGCCGCCIG GGIGGCGCIG GCGGAGGGAC 600 TGGGTGTGGC CGTGTACGCA GCGGCTGTGC TGCTGGGTGC TGGCTGTGCC ACCATCCTCG 660 TCACCTCGCT GGCCATGACG GCCGACCTCA TCGGTCCCCA CACGAACAGC GGACTKTCGT 720 30 GTACGGCTCC ATGAGCTTCT TGGATAAGGT GGCCAATGGG CTGGCAGTCA TGGCCATCCA 780 GAGCCTGCAC CCTTGCCCCT CAGAGCTCTG CTGCAGGGCC TGCGTGAGCT TTTACCACTG 840 35 GGCGATGGTG GCTGTGACGG GCGGCGTGGG CGTGGCCGCT GCCCTGTGTC TCTGTAGCCT 900 960 CCTGCTGTGG CCGACCCGCC TGCGACGCTG GGACCGTGAT GCCCGGCCCT GACTCCTGAC AGCCTCCTGC ACCTGTGCAA GGGAACTGTG GGGACGCACG AGGATGCCCC CCARGGCCTT 1020 40 GGGGAAAAGC CCCCACTGCC CCTCACTCTT CTCTGGACCC CCACCCTCCA TCCTCACCCA 1080 GCTCCCGGGG GTGGGGTCGG GTGAGGGCAG CAGGGATGCC CGCCAGGGAC TTGCAAGGAC 1140 45 CCCCTGGGTT TTGAGGGTGT CCCATTCTCA ACTCTAATCC ATCCCAGCCC TCTGGAGGAT 1200 TTGGGGTGCC CCTCTCGGCA GGGAACAGGA AGTAGGAATC CCAGAAGGGT CTGGGGGAAC 1260 CCTAACCCTG AGCTCAGTCC AGTTCACCCC TCACCTCCAG CCTGGGGGTC TCCAGACACT 1320 50 GCCAGGGCCC CCTCAGGACG GCTGGAGCCT GGAGGAGACA GCCACGGGGT GGTGGGCTGG 1380 GCCTGGACCC CACCGTGGTG GGCAGCAGGG CTGCCCGGCA GGCTTGGTGG ACTCTGCTGG 1440 55 1500 1525 AAAAAAAAA AAACCCACCG TCCGC

(2) INFORMATION FOR SEQ ID NO: 80:

5 10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 1563 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 80:	
	AATTCGGCAC GAGNCAGAAA CCTGCGGAAA ATGGTAGCGA TGGCGGCTGG GCCGAGTGGG	60
15	TGTCTGGTGC CGGCGTTTGG GCTACGGTTG TTGTTGGCGA CTGTGCTTCA AGCGGTGTCT	120
	GCTTTTGGGG CAGAGTTTTC ATCGGAGGCA TGCAGAGAGT TAGGCTTTTC TAGCAACTTG	180
20	CTTTGCAGCT CTTGTGATCT TCTCGGACAG TTCAACCTGC TTCAGCTGGA TCCTGATTGC	240
	AGAGGATGCT GTCAGGAGGA AGCACAATTT GAAACCAAAA AGCTGTATGC AGGAGCTATT	300
	CTTGAAGTTT GTGGATGAAA ATTGGGAAGG TTCCCTCAAG TCCAAGCTTT TGTTAGGAGT	360
25	GATAAACCCA AACTGTTCAG AGGACTGCAA ATCAAGTATG TCCGTGGTTC AGACCCTGTA	420
	TTAAAGCTTT TGGACGACAA TGGGAACATT GCTGAAGAAC TGAGCATTCT CAAATGGAAC	480
30	ACAGACAGTG TAGAAGAATT CCTGAGTGAA AAGTTGGAAC GCATATAAAT CTTGCTTAAA	540
	TTTTGTCCTA TCCTTTTGTT ACCTTATCAA ATGAAATATT ACAGCACCTA GAAAATAATT	600
	TAGTTTTGCT TGCTTCCATT GATCAGTCTT TTACTTGAGG CATTAAATAT CTAATTAAAT	660
35	CGTGAAATGG CAGTATAGTC CATGATATCT AAGGAGTTGG CAAGCTTAAC AAAACCCATT	720
	TTTTATAAAT GTCCATCCTC CTGCATTTGT TGATACCACT AACAAAATGC TTTGTAACAG	780
40	ACTTGCGGTT AATTATGCAA ATGATAGTTT GTGATAATTG GTCCAGTTTT ACGAACAACA	840
	GATTTCTAAA TTAGAGAGGT TAACAAGACA GATGATTACT ATGCCTCATG TGCTGTGTGC	900
	TCTTTGAAAG GAATGACAGC AGACTACAAA GCAAATAAGA TATACTGAGC CTCAACAGAT	960
45	TGCCTGCTCC TCAGAGTCTC TCCTATTTTT GTATTACCCA GCTTTCTTTT TAATACAAAT	1020
	GTTATTTATA GTTTACAATG AATGCACTGC ATAAAAACTT TGTAGCTTCA TTATTGTAAA	1080
50	ACATATTCAA GATCCTACAG TAAGAGTGAA ACATTCACAA AGATTTGCGT TAATGAAGAC	1140
	TACACAGAAA ACCTTTCTAG GGATTTGTGT GGATCAGATA CATACTTGGC AAATTTTTIGA	1200
	GTTTTACATT CTTACAGAAA AGTCCATTTA AAAGTGATCA TTTGTAAGAC CAAAATATAA	1260
55	ATAAAAAGTT TCAAAAAATCT ATCTGAATTT GGAATTCTTC TGGTTTGTTC TTTCATGTTT	1320
	AAAAATGATG TTTTTCAATG CATTTTTTC ATGTAAGCCC TTTTTTTAGC CAAAATGTAA	1380
60	AAATGGCTGT AATATTTAAA ACTTATAACA TCTTATTGTT GGTAATAGTG CTTTATATTT	1440

	GTCTGATTTT	ATTTTTCAAA	GTTTTTTCAT	TTATGAACAC	ATTTTCATTG	GTATATTATT	1500
	TAAGGAATAT	CTCTTGATAT	AGAATTTTTA	TATTAAAAAT	GATTTTTCTT	TGCTTAAAAA	1560
5	AAA						1563

10 (2) INFORMATION FOR SEQ ID NO: 81:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1020 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 81:

20	TGCACGCTGG	CCATGTGGGN	GTTGGGCCAC	TGCGACCCCC	GGCGCTGCAC	GGGCCGCAAG	60
	CIGGCCCGCC	TGGGGCTGGT	GCGCTGCCTG	CGCCTGGGCC	ACAGATTCGG	CGGTCTGGTG	120
25	CTGAGCCCCG	TGGGCAAGCA	GTACGCGTCC	CCCGCAGACA	GACAGCTGGT	GGCGCAGTCT	180
23	GGGTCGCCG	TCATCGACTG	CTCCTGGGCC	AGGCTGGACG	AGACACCGTT	TGGGAAGATG	240
	CGAGGGAGCC	ACTTGCGCCT	GTTGCCCTAC	CTGGTGGCCG	CCAACCCCGT	GAACTATGGC	300
30	CGGCCCTACA	GACTTTCCTG	CGTGGAAGCG	TTTGCTGCCA	CCTTCTGCAT	CGTAGGCTTT	360
	CCAGACCTTG	CTGTCATTTT	GCTGCGGAAG	TTTAAATGGG	GCAAGGGCTT	CTTGGACCTG	420
35	AACCGCCAGC	TCCTGGACAA	GTACGCGGCC	TGCGGCAGCC	CGGAGGAGGT	GCTGCAGGCG	480
33	GAGCAGGAGT	TCTTGGCCAA	TGCCAAGGAG	AGCCCCCAGG	AGGAGGAGAT	CGATCCCTTC	540
	GATGTGGATT	CAGGGAGAGA	GTTTGGAAAC	CCCAACAGGC	CTGTGGCCAG	CACCCGGCTG	600
40	CCCTCGGACA	CTGATGACAG	TGATGCGTCT	' GAGGACCCAG	GGCCTKGCGC	CGAGCGCGGA	660
	GGAGCCAGCA	GCAGCTGCTG	TGAAGAGGAG	CAGACGCAGG	GACGGGGGC	TGAGGCCAGG	720
15	GCCCCGGCTG	AGGTTTGGAA	AGGAATCAAG	AAACGCCAGA	GAGACTGAGG	GTTGCAGACA	780
45	CATATATTT	TGAGGCTGGG	; TGACGAGAAA	ATCTAGAGAC	ATGAGGGACA	TAAATGGGCC	840
	TGGCAGCCTC	GGCTCTTTGC	GGCTGCTGGC	C AGGACTGAGC	TGTCCGGGTT	CTCCCCACAC	900
50	TTCCAGCACA	A GCTGTGCTC1	GTGTCCTGC	TCGGCGCTCT	CGCAAATGAA	GCTGCAGGCC	960
	AAGAAAAAA	AAAAAAAA	AAAAAAAA	AAAAAAAA	AAAAAAAAA	GGGGGGGGC	1020

55

(2) INFORMATION FOR SEQ ID NO: 82:

(i) SEQUENCE CHARACTERISTICS:

60 (A) LENGTH: 770 base pairs

(B) TYPE: nucleic acid

	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 82:	
	TCGACCCACG CGTCCGGGCC GCCGTAGCGC GTCTTGGGTC TCCCGGCTGC CGCTGCTGCC	60
10	GCCGCCGCCT CGGGTCGTGG AGCCAGGAGC GACGTCACCG CCATGGCAGG CATCAAAGCT	120
10	TTGATTAGTT TGTCCTTTGG AGGAGCAATC GGACTGATGT TTTTRATGCT TGGATGTGCC	180
	CTTCCAATAT ACAACAAATA CTGGCCCCTC TTTGTTCTAT TTTTTTACAT CCTTTCACCT	240
15	ATTCCATACT GCATAGCAAG AAGATTAGTG GATGATACAG ATGCTATGAG TAACGCTTGT	300
	AAGGAACTTG CCATCTTTCT TACAACGGGC ATTGTCGTGT CAGCTTTTGG ACTCCCTATT	360
20	GTATTTGCCA GAGCACATCT GATTGAGTGG GGAGCTTGTG CACTTGTTCT CACAGGAAAC	420
20	ACAGTCATCT TTGCAACTAT ACTAGGCTTT TTCTTGGTCT TTGGAAGCAA TGACGACTTC	480
	AGCTGGCAGC AGTGGTGAAA AGAAATTACT GAACTATTGT CAAATGGACT TCCTGTCATT	540
25	TGTTGGCCAT TCACGCACAC AGGAGATGGG GCAGTTAATG CTGAATGGTA TAGCAAGCCT	600
	CTTGGGGGTA TITTAGGTGC TCCCTTCTCA CTTTTATTGT AAGCATACTA TTTTCACAGA	660
30	GACTTGCTGA AGGATTAAAA GGATTTTCTC TTTTGGAAAA AAAAAAAAA AAAAACYCGA	720
30	GGGGGGGCCC GTWCCCATTC SCCCYATATG AATTCCNTTT TTACAATCCC	770
35	(2) INFORMATION FOR SEQ ID NO: 83:	
40	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 481 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
45	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 83:	
	GAATTCGGCA CGAGCATAGT GTTAACCACT AGAATTCACT GCCCTTCCTA TCCAAAAATG	60
	ACACTACTGA TCATTTTTCT TCCTTTTSCT TTTACAACAT TMACAAATTC AGGTGGCTCT	120
50	TTCCCAGTAC GGTAGGCTGA TTCGTATGGA TGCACCACGG TTGGTGACTC CCCCCACCCC	180
	ACAGAGTTTC TGGCGTTCAT TCGGTTGAAC CCAAGGCCAG CAAGGGCTGA CTGGGAACAA	240
55	ACCGAACACT AGGCCGTGAA CCAATCGTCT CTCCGTGCCC GGGAGCGAMC CCGGGGGCCT	300
	TTCACTCTCC CAAGGACTCC ANGGGGGGGC CGGGTACCCA ATTCCGCCCC TATAGTGAAT	360
	CCGTNATTAC AATTCCACNT GGGCCGTCCN TTTTTACAAA CGTTCCGTTG AACTGGGAAA	420
60	AACCCCTTGG CGGTTTACCC CAACTTTAAT CCGCCTTTGC AAGCACATCC CCCCCCTTTT	480

	c ·	481
5		
J	(2) INFORMATION FOR SEQ ID NO: 84:	
10	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 644 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 84:	
	GCTGGGATAG AGCATGAAAG GAGAACTGCT CCCTTTTCTG TTTCTCACAG TTTGGTTATG	60
•	GCTTTATAAA CTTKTATTTG GTGAAAGCCC CAGATACCCA AATGTCATTG GCAAAACTTA	120
20	TTTTTTTTC TGGACAGATC AGATTCTAG AGAGAGCAGA TTTCTAGAGA GATTAGCATT	180
	CATAGTAAGT GAAAATTGTC TAATTTTTTT AATCCATGCT ATTACTGGGC AGTAGGTCTA	240
25	ATTTTTTTT ACAAAAAATA GATCTATTTT CCTTATATAT TGATTTAGAA TCTTAAGTTA	300
	GAATTTTATA GAAGAAATGT CTGAGCAGTT CTATGTATGG AGGAGCAATT CAGCTTTTCA	360
20	GCAGCAACTT TATCTTTTGC CACTAGAGGG AGATCTGTGG TTGCTTTCTC CTTTGGAGAA	420
30	TAGCTGCTTT GCTTTTATTT TTAATTTCTA AGGTTGGAAT AGAACTTATT CTCAAAATTC	480
	CTTTAGTGTT ATTAAATATT TTCATTTATT AGTCAAAGGT AAGTTAATTA AGCTTGTTTA	540
35	ATGATGCCAA TCTTATGCTT TTCTGTAATC TTCAATTTTT AATAAATGTG AGTTAGATAC	600
	TAAGTGAAAA AAAAAAAAA AAAAAAAAA AAAAAAAAA	644
40		
	(2) INFORMATION FOR SEQ ID NO: 85:	
45	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1351 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 85:	
	GCCACGAGTG CGCASGCGTG GGGCTCTCTC CTTGTCAGTC GGCGCCGCGT GCGGGCTGGT	60
<i>e e</i>	GGCTCTGTGG CAGCGGCGGC GGCAGGACTC CGGCACTATG AGCGGCTTCA GCACCGAGGA	120
55	GCGCGCCGCG CCNTTCTCCC TGGAGTACCG AGTCTTCCTC AAAAATGAGA AAGGACAATA	180
	TATATCTCCA TTTCATGATA TTCCAATTTA TGCAGATAAG GATGTGTTTC ACATGGTAGT	240
60	TGAAGTACCA CGCTGGTCTA ATGCAAAAAT GGAGATTGCT ACAAAGGACC CTTTAAACCC	300

	TATTAAACAA GATGTGAAAA AAGGAAAACT TCGCTATGTT GCGAATTTGT TCCCGTATAA	360
5	AGGATATATC TGGAACTATG GTGCCATCCC TCAGACTTGG GAAGACCCAG GGCACAATGA	420
J	TAAACATACT GGCTGTTGTG GTGACAATGA CCCAATTGAT GTGTGTGAAA TTGGAAGCAA	480
	GGTATGTGCA AGAGGTGAAA TAATTGGCGT GAAAGTTCTA GGCATATTGG CTATGATTGA	540
10	CGAAGGGGAA ACCGACTGGA AAGTCATTGC CATTAATGTG GATGATCCTG ATGCAGCCAA	600
	TTATAATGAT ATCAATGATG TCAAACGGCT GAAACCTGGC TACTTAGAAG CTACTGTGGA	660
15	CTGGTTTAGA AGGTATAAGG TTCCTGATGG AAAACCAGAA AATGAGTTTG CGTTTAATGC	720
13	AGAATTTAAA GATAAGGACT TTGCCATTGA TATTATTAAA AGCACTCATG ACCATTGGAA	780
	AGCATTAGTG ACTAAGAAAA CGAATGGAAA AGGAATCAGT TGCATGAATA CAACTTTGTC	840
20	TGAGAGCCCC TTCAAGTGTG ATCCTGATGC TGCCAGAGCC ATTGTGGATG CTTTACCACC	900
	ACCCTGTGAA TCTGCCTGCA CAGTACCAAC AGACGTGGAT AAGTGGTTCC ATCACCAGAA	960
25	AAACTAATGA GATTTCTCTG GAATACAAGC TGATATTGCT ACATCGTGTT CATCTGGATG	1020
	TATTAGAAGT AAAAGTAGTA GCTTTCAAA GCTTTAAATT TGTAGAACTC ATCTAACTAA	1080
	AGTAAATTCT GCTGTGACTA ATCCAATATA CTCAGAATGT TATCCATCTA AAGCATTTTT	1140
30	CATATCTCAA CTAAGATAAC TTTTAGCACA TGCTTAAATA TCAAAGCAGT TGTCATTTGG	1200
	AAGTCACTTG TGAATAGATG TGCAAGGGGA GCACATATTG GATGTATATG TTACCATATG	1260
35	TTAGGAAATA AAATTATTTT GCTGAAAAAA AAAAAAAAA AACCNCGGG GGGGCCCCGG	1320
	TCCCCATTIG GCCCTTTGGG GGGNGGTTTT A	1351
40	(2) INFORMATION FOR SEQ ID NO: 86:	
	(i) SEQUENCE CHARACTERISTICS:	
45	(A) LENGTH: 2527 base pairs (B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 86:	
50		
	CTCTTGCTAC CTTCCCGGCG CAGAGAACCC CGGCTGCTCA GCGCGCTCCG GGGTCATGGA	60
55	GATCCCCGGG AGCCTGTGCA AGAAAGTCAA GCTGAGCAAT AACGCGCAGA ACTGGGGAAT	120
ر ر	GCAGAGAGCA ACCAATGTCA CCTACCAAGC CCATCATGTC AGCAGGAACA AGAGAGGTCA	180
	GGTGGTGGGG ACCAGAGGTG GCTTTCGTGG TTGCACAGTT TGGCTAACAG GCTTGTCTGG	240
50	AGCGGGAAAG ACTACTGTGA GCATGGCCTT GGAGGAGTAC CTGGTTTGTC ATGGTATTCC	300

	ATGCTACACT CTGGATGGTG ACAATATTCG TCAAGGTCTC AATAAAAATC TTGGCTTTAG	360
	TCCTGAAGAC AGAGAAGAGA ATGTTCGACG CATCGCAGAA GTTGCTAAAC TGTTTGCAGA	420
5	TGCTGGCTTA GTGTGCATCA CAAGTTTCAT ATCACCTTAC ACTCAGGATC GCAACAATGC	480
	AAGGCAAATT CATGAAGGTG CAAGTTTACC GTTTTTTGAA GTATTTGTTG ATGCTCCTCT	540
10	GCATGTTTGT GAACAGAGGG ATGTCAAAGG ACTCTACAAA AAAGCCCGGG CAGGAGAAAT	600
10	TAAAGGTTTC ACTGGGATCG ATTCTGAATA TGAAAAGCCA GAGGCCCCTG AGTTGGTGCT	660
	GAAAACAGAC TCCTGTGATG TAAATGACTG TGTCCAGCAA GTTGTGGAAC TTCTACAGGA	720
15	ACGGGATATT GTACCTGTGG ATGCATCTTA TGAAGTAAAA GAACTATATG TGCCAGAAAA	780
	TAAACTTCAT TTGGCAAAAA CAGATGCGGA AACATTACCA GCACTGAAAA TTAATAAAGT	840
20	GGATATGCAG TGGGTGCAGG TTTTGGCAGA AGGTTGGGCA ACCCCATTGA ATGGCTTTAT	900
20	GAGAGAGAG GAGTACTTGC AGTGCCTTCA TTTTGATTGT CTTCTGGATG GAGGTGTCAT	960
	TAACTTGTCA GTACCTATAG TTCTGACTGC GACTCATGAA GATAAAGAGA GGCTGGACGG	1020
25	CTGTACAGCA TTTGCTCTGA TGTATGAGGG CCGCCGTGTG GCCATTCTTC GCAATCCAGA	1080
	GTTTTTTGAG CACAGGAAAG AGGAGCGCTG TGCCAGACAG TGGGGAACGA CATGCAAGAA	1140
30	CCACCCTAT ATTAAGATGG TGATGGAACA AGGAGATTGG CTGATTGGAG GAGATCTTCA	1200
30	AGTCTTGGAT CGAGTTTATT GGAATGATGG TCTTGATCAG TATCGTCTTA CTCCTACTGA	1260
	GCTAAAGCAG AAATTTAAAG ATATGAATGC TGATGCTGTC TTTGCATTTC AACTACGCAA	1320
35	CCCAGTGCAC AATGGACATG CCCTGTTAAT GCAGGATACC CATAAGCAAC TTCTAGAGAG	1380
	GGGCTACCGG CGCCCTGTCC TCCTCCTCCA CCCTCTGGGT GGCTGGACAA AGGATGACGA	1440
40	TGTTCCTTTG ATGTGGCGTA TGAAGCAGCA TGCTGCAGTG TTGGAGGAAG GAGTTCTGAA	1500
10	TCCTGAGACG ACAGTGGTGG CCATCTTCCC ATCTCCCATG ATGTATGCTG GACCAACTGA	1560
	GGTCCAGTGG CATTGCAGAG CACGGATGGT TGCAGGAGCC AACTTTTACA TTGTTGGACG	1620
45	AGACCCTGCT GGCATGCCTC ATCCAGAAAC AGGGAAGGAT CTTTATGAGC CAAGTCATGG	1680
	TGCCAAAGTG CTGACGATGG CCCCTGGTTT AATCACTTTG GAAATAGTTC CCTTTCGAGT	1740
50	TGCAGCTTAC AACAAGAAAA AGAAGCGTAT GGACTACTAT GACTCTGAAC ACCATGAAGA	1800
50	CTTTGAATTT ATTTCAGGAA CACGAATGCG CAAACTTGCT CGAGAAGGCC AGAAACCACC	1860
	TGAAGGTTTC ATGGCTCCCA AGGCTTGGAC CGTGCTGACA GAATACTACA AATCCTTGGA	1920
55	GAAAGCTTAG GCTGTTAACC CAGTCACTCC ACCTTTGACA CATTACTAGT AACAAGAGGG	1980
	GACCACATAG TCTCTGTTGG CATTTCTTTG TGGTGTCTGT CTGGACATGC TTCCTAAAAA	2040
60	CAGACCATTT TCCTTAACTT GCATCAGTTT TGGTCTGCCT TATGAGTTCT GTTTTGAACA	2100

	AGTGTAACAC ACTGATGGTT TTAATGTATC TTTTCCACTT ATTATAGTTA TATTCCTACA	2160
	ATACAATTTT AAAATTGTCT TTTTATATTA TATTTATGCT TCTGTGTCAT GATTTTTTCA	2220
5	AGCTGTTATA TTAGTTGTAA CCAGTAGTAT TCACATTAAA TCTTGCTTTT TTTCCCCTTA	2280
	AAAAAAGAAA AAAATTACCA AACAATAAAC TTGGCTAGAC CTTGTTTTGA GGATTTTACA	2340
10	AGACCTTTGT AGCGATTAGA TTTTTTTTCT ACATTGAAAA TAGAAACTGC TTCCTTTCTT	2400
10	CTTTCCAGTC AGCTATTGGT CTTTCCAGCT GTTATAATCT AAAGTATTCT TATGATCTGT	2460
	GTAAGCTCTG AATGAACTTC TTTACTCAAT AAAATTAATT TTTTGGCTTC TTAAAAAAAA	2520
15	AAAAAA	2527
20	(2) INFORMATION FOR SEQ ID NO: 87:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 2566 base pairs (B) TYPE: nucleic acid	
25	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 87:	
30	CCCAAGAATT CGGCACGAGC GNGGCAWAAK TGGGATTTCT GAAACCTGTA GGCCCCAAGC	-60
	CCATCAACTT GCCCAAAGAA GATTCCAAAC CTACATTTCC CTGGCCTSCT GGAAACAAGC	120
	CATCTCTTCA CAGTGTAAAC CAAGACCATG ACTTAAAGCC ACTAGGCCGA AATCTGGGCC	180
35	TACTCCTCCA ACCTCAGAAA ATGAACAGAA GCAAGCKTTT CCCAAATTGA CTGGGGTTAA	240
	AGGGAAATTT ATGTCAGCAT CACAAGATCT TGAACCCAAG CCCCTCTTCC CCAAACCCGC	300
40	CTTTGGCCAG AAGCCGCCCC TAAGTACCGA GAACTCCCAT GAAGACGAAA GCCCCATGAA	360
	GAATGTGTCT TCATCAAAAG GGTCCCCAGC TCCCCTGGGA GTCAGGTCCA AAAGCGGCCC	420
45	TTTAAAACCA GCAAGGGAAG ACTCAGAAAA TAAAGACCAT GCAGGGGAGA TTTCAAGTTT	480
45	GCCCTTTCCT GGAGTGGTTT TGAAACCTGC TGCGAGCAGG GGAGGCCCAG GTCTCTCCAA	540
	AAATGGTGAA GAAAAAAAGG AAGATAGGAA GATAGATGCT GCTAAGAACA CCTTCCAGAG	600
50	CAAAATAAAT CAGGAAGAGT TGGCCTCAGG GACTCCTCCT GCCAGGTTCC CTAAGGCCCC	660
	TTCTAAGCTG ACAGTGGGG GGCCATGGGG CCAAAGTCAG GAAAAGGAAA AGGGAGACAA	720
55	GAATTCAGCC ACCCCGAAAC AGAAGCCATT GCCTCCCTTG TTTACCTTGG GTCCACCTCC	780
55	ACCAAAACCC AACAGACCAC CAAATGTTGA CCTGACGAAA TTCCACAAAA CCTCTTCTGG	840
	AAACAGTACT AGCAAAGGCC AGACGTCTTA CTCAACAACT TCCCTGCCAC CACCTCCACC	900
60	ATCCCATCCG GCCAGCCAAC CACCATTGCC AGCATCTCAC CCATCACAAC CACCAGTCCC	960

WO 98/42738

PCT/US98/05311

	AAGCCTACCT CCCAGAAACA TTAAACCTCC GTTTGACCTA AAAAGCCCTG TCAATGAAGA	1020
_	CAATCAAGAT GGTGTCACGC ACTCTGATGG TGCTGGAAAT CTAGATGAGG AACAAGACAG	1080
5	TGAAGGAGAA ACATATGAAG ACATAGAAGC ATCCAAAGAA AGAGAGAAGA AAAGGGAAAA	1140
	GGAAGAAAAG AAGAGGTTAG AGCTGGAGAA AAAGGAACAG AAAGAGAAAG AAAAGAAAGA	1200
10	ACAAGAAATA AAGAAGAAAT TTAAACTAAC AGGCCCTATT CAAGTCATCC ATCTTGCAAA	1260
	AGCTTGTTGT GATGTCAAAG GAGGAAAGAA TGAACTGAGC TTCAAGCAAG GAGAGCAAAT	1320
1.5	TGAAATCATC CGCATCACAG ACAACCCAGA AGGAAAATGG TTGGGCAGAA CAGCAAGGGG	1380
15	TTCATATGGC TATATTAAAA CAACTGCTGT AGAGATTGAC TATGATTCTT TGAAACTGAA	1440
	AAAAGACTCT CTTGGTGCCC CTTCAAGACC TATTGAAGAT GACCAAGAAG TATATGATGA	1500
20	TGTTGCAGAG CAGGATGATA TTAGCAGCCA CAGTCAGAGT GGAAGTGGAG GGATATTCCC	1560
	TCCACCACCA GATGATGACA TTTATGATGG GATTGAAGAG GAAGATGCTG ATGATGGCTC	1620
25	CACACTACAG GTTCAAGAGA AGAGTAATAC GTGGTCCTGG GGGATTTTGA AGATGTTAAA	1680
23	GGGAAAAGAT GACAGAAAGA AAAGTATACG AGAGAAACCT AAAGTCTCTG ACTCAGACAA	1740
	TAATGAAGGT TCATCTTTCC CTGCTCCTCC TAAACAATTG GACATGGGAG ATGAAGTTTA	1800
30	CGATGATGTG GATACCTCTG ATTTCCCTGT TTCATCAGCA GAGATGAGTC AAGGAACTAA	1860
	TGTTGGAAAA GCTAAGACAG AAGAAAAGGA CCTTAAGAAG CTAAAAAAAGC AGRAAAAARA	1920
35	ARAAAAAGAC TTCAGGAAAA AATTTAAATA TGATGGTGAA ATTAGAGTCC TATATTCAAC	1980
33	TAAAGTTACA ACTTCCATAA CTTCTAAAAA GTGGGGAACC AGAGATCTAC AGGTAAAACC	2040
	TGGTGAATCT CTAGAAGTTA TACAAACCAC AGATGACACA AAAGTTCTCT GCAGAAATGA	2100
40	AGAAGGGAAA TATGGTTATG TCCTTCGGAG TTACCTAGCG GACAATGATG GAGAGATCTA	2160
	TGATGATATT GCTGATGGCT GCATCTATGA CAATGACTAG CACTCAACTT TGGTCATTCT	2220
45	GCTGTGTTCA TTAGGTGCCA ATGTGAAGTC TGGATTTTAA TTGGCATGTT ATTGGGTATC	2280
43	AAGAAAATTA ATGCACAAAA CCACTTATTA TCATTTGTTA TGAAATCCCA ATTATCTTTA	2340
	CAAAGTGTTT AAAGTTTGAA CATAGAAAAT AATCTCTCTG CTTAATTGTT ATCTCAGAAG	2400
50	ACTACATTAG TGAGATGTAA GAATTATTAA ATATTCCATT TCCGCTTTGG CTACAATTAT	2460
	GAAGAAGTTG AAGGTACTTC TTTTAGACCA CCAGTAAATA ATCCTCCTTC AAAAAATAAA	2520
E	AATAAAAAAA AAAAAAAAA ACTCGAGGGG GGGCCCGGTA CCCAAT	2566
55		

(2) INFORMATION FOR SEQ ID NO: 88:

5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 540 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 88:	
10	GAATTCGGCA CGAGGCTTTC TGTGTCCTCT GTGGCTGCTT TAGTGTGCCA CCAGGGGCAG	60
	ACTTGGGTGG GTTGCAGCAG AGATGGCATG GCCCTCAAGG TCCAAGATGT TTACTCTCTT	120
	GCCGGTCCTC TGTTATCTCT GGTCTTTGTG GTTGCCACAG TTTTCTTGGA TCCAGGAGTT	180
15	AAAGGCAGTC CTGAGGGATG ATGGCCTCAT CTCCGCAGTT GCYTGGAATG CTGAATTTCA	240
	GACGTGCTAA AGGAGGGTTG CAGACATTGT GTGGWATGCA TTCAGACCCC AGATGTGGGT	300
20	GCAGGAAGGC AGGCATGGCA CAGCCAGGTA GAGACTGGTT TCCAGGCCCA AGCAGCCTTC	360
	AGCAGCTGTG CGCCTTGTTT CTGATGTTGT TTGGGAGTAA GAATAATGTA GACATGGGGG	420
	GTCATGARGC TCAATAAAAA CTTCAAGGAA ACCTCCCATG GCATGGTTGG GCGCAGTGAC	480
25	TCATGCCTGT AACCCCAGCA CTGTGGAATG CCAAGGTGGA AGGATCGCTT GAGGCCAAGA	540
30 35	(2) INFORMATION FOR SEQ ID NO: 89: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1863 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 89:	
40	TCGACCCACG CGTCCGGCGA GATCCCTACC GCAGTAGCCG CCTCTGCCGC CGCGGAGCTT	60
15	CCCGAACCTC TTCAGCCGCC CGGAGCCGCT CCCGGAGCCC GGCCGTAGAG GCTGCAATCG	120
	CCCGAACCTC TTCAGCCGCC CGGAGCCGCT CCCGGAGCCC GGCCGTAGAG GCTGCAATCG CAGCCGGGAG CCCGCAGGCC GCGCCCGAG CCCTTCGAGG GCGCCCCAGG	120 180
45		
43	CAGCCGGGAG CCCGCAGCCC GCGCCCCGAG CCCGCCGCCG CCCTTCGAGG GCGCCCCAGG	180
50	CAGCCGGGAG CCCGCAGCCC GCGCCCCGAG CCCGCCGCCG CCCTTCGAGG GCGCCCCAGG CCGCGCCCATG GTGAAGGTGA CGTTCAACTC CGCTCTGGCC CAGAAGGAGG CCAAGAAGGA	180 240
	CAGCCGGGAG CCCGCAGCCC GCGCCCCGAG CCCGCCGCCG CCCTTCGAGG GCGCCCCAGG CCGCGCCATG GTGAAGGTGA CGTTCAACTC CGCTCTGGCC CAGAAGGAGG CCAAGAAGGA CGAGCCCAAG AGCGGCGAGG AGGCGCTCAT CATCCCCCCC GACGCCGTCG CGGTGGACTG	180 240 300
50	CAGCCGGGAG CCCGCAGCCC GCGCCCCGAG CCCGCCGCCG CCCTTCGAGG GCGCCCCAGG CCGCGCCATG GTGAAGGTGA CGTTCAACTC CGCTCTGGCC CAGAAGGAGG CCAAGAAGGA CGAGCCCAAG AGCGGCGAGG AGGCGCTCAT CATCCCCCCC GACGCCGTCG CGGTGGACTG CAAGGACCCA GATGATGTGG TACCAGTTGG CCAAAGAAGA GCCTGGTGTT GGTGCATGTG	180 240 300 360
	CAGCCGGGAG CCCGCAGCCC GCGCCCCGAG CCCGCCGCCG CCCTTCGAGG GCGCCCCAGG CCGCGCCATG GTGAAGGTGA CGTTCAACTC CGCTCTGGCC CAGAAGGAGG CCAAGAAGGA CGAGCCCAAG AGCGGCGAGG AGGCGCTCAT CATCCCCCCC GACGCCGTCG CGGTGGACTG CAAGGACCCA GATGATGTGG TACCAGTTGG CCAAAGAAGA GCCTGGTGTT GGTGCATGTG CTTTGGACTA GCATTTATGC TTGCAGGTGT TATTCTAGGA GGAGCATACT TGTACAAATA	180 240 300 360 420
50	CAGCCGGGAG CCCGCAGCCC GCGCCCCGAG CCCGCCGCG CCCTTCGAGG GCGCCCCAGG CCGCGCCATG GTGAAGGTGA CGTTCAACTC CGCTCTGGCC CAGAAGGAGG CCAAGAAGGA CGAGCCCAAG AGCGGCGAGG AGGCGCTCAT CATCCCCCCC GACGCCGTCG CGGTGGACTG CAAGGACCCA GATGATGTGG TACCAGTTGG CCAAAGAAGA GCCTGGTGTT GGTGCATGTG CTTTGGACTA GCATTTATGC TTGCAGGTGT TATTCTAGGA GGAGCATACT TGTACAAATA TTTTGCACTT CAACCAGATG ACGTGTACTA CTGTGGAATA AAGTACATCA AAGATGATGT	180 240 300 360 420 480

	TCTTAACCTG GATAAGTGCT ATGTGATCCC TCTGAACACT TCCATTGTTA TGCCACCCAG	720
_	AAACCTACTG GAGTTACTTA TTAACATCAA GGCTGGAACC TATTTGCCTC AGTCCTATCT	780
5	GATTCATGAG CACATGGTTA TTACTGATCG CATTGAAAAC ATTGATCACC TGGGTTTCTT	840
	TATTTATCGA CTGTGTCATG ACAAGGAAAC TTACAAACTG CAACGCAGAG AAACTATTAA	900
10	AGGTATTCAG AAACGTGAAG CCAGCAATTG TTTCGCAATT CGGCATTTTG AAAACAAATT	960
	TGCCGTGGAA ACTTTAATTT GTTCTTGAAC AGTCAAGAAA AACATTATTG AGGAAAATTA	1020
1.5	ATATCACAGC ATAACCCCAC CCTTTACATT TTGTGCAGTG ATTATTTTTT AAAGTCTTCT	1080
15	TTCATGTAAG TAGCAAACAG GGCTTTACTA TCTTTTCATC TCATTAATTC AATTAAAACC	1140
	ATTACCTTAA AATTTTTTC TITCGAAGTG TGGTGTCTTT TATATTTGAA TTAGTAACTG	1200
20	TATGAAGTCA TAGATAATAG TACATGTCAC CTTAGGTAGT AGGAAGAATT ACAATTTCTT	1260
	TAAATCATTT ATCTGGATTT TTATGTTTTA TTAGCATTTT CAAGAAGACG GATTATCTAG	1320
25	AGAATAATCA TATATATGCA TACGTAAAAA TGGACCACAG TGACTTATTT GTAGTTGTTA	1380
25	GTTGCCCTGC TACCTAGTTT GTTAGTGCAT TTGAGCACAC ATTTTAATTT TCCTCTAATT	1440
	AAAATGTGCA GTATTTTCAG TGTCAAATAT ATTTAACTAT TTAGAGAATG ATTTCCACCT	1500
30	TTATGTTTTA ATATCCTAGG CATCTGCTGT AATAATATTT TAGAAAATGT TTGGAATTTA	1560
	AGAAATAACT TGTGTTACTA ATTTGTATAA CCCATATCTG TGCAATGGAA TATAAATATC	1620
25	ACAAAGTTGT TTAACTAGAC TGCGTGTTGT TTTTCCCGTA TAATAAAACC AAAGAATAGT	1680
35	TTGGTTCTTC AAATCTTAAG AGAATCCACA TAAAAGAAGA AACTATTTTT TAAAAAATTCA	1740
	CTTCTATATA TACAATGAGT AAAATCACAG ATTTTTTCTT TAAATAAAAA TAAGTCATTT	1800
40	TAATAACTAA ACCAGATTCT TTGTGATACT ATTAANGTAA CATTTAGCCC CAAAAAAAAA	1860
	AAA	1863
15		
45	(2) INFORMATION FOR SEQ ID NO: 90:	
	(2) INFORMATION FOR SEQ ID NO. 30. (i) SEQUENCE CHARACTERISTICS:	
50	(A) LENGTH: 2478 base pairs (B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
55	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 90:	
	GGCACAGCGG CACGAGGTGA GCTGAGCCGG TGGGTGAGCG GCGGCCACGG CATCCTGTGC	60
	TGTGGGGGCT ACGAGGAAAG ATCTAATTAT CATGGACCTG CGACAGTTTC TTATGTGCCT	120

	GTCCCTGTGC	ACAGCCTTTG	CCTTGAGCAA	ACCCACAGAA	AAGAAGGACC	GTGTACATCA	180
	TGAGCCTCAG	CTCAGTGACA	AGGTTCACAA	TGATGCTCAG	AGTTTTGATT	ATGACCATGA	240
5	TGCCTTCTTG	GGTGCTGAAG	AAGCAAAGAC	CTTTGATCAG	CTGACACCAG	AAGAGAGCAA	300
	GGAAAGGCTT	GGAAAGATTG	TAAGTAAAAT	AGATGGCGAC	AAGGACGGGT	TTGTCACTGT	360
10	GGATGAGCTC	AAAGACTGGA	TTAAATTTGC	ACAAAAGCGC	TGGATTTACG	AGGATGTAGA	420
10	GCGACAGTGG	AAGGGGCATG	ACCTCAATGA	GGACGGCCTC	GTTTCCTGGG	AGGAGTATAA	480
	AAATGCCACC	TACGGCTACG	TTTTAGATGA	TCCAGATCCT	GATGATGGAT	TTAACTATAA	540
15	ACAGATGATG	GTTAGAGATG	AGCGGAGGTT	TAAAATGGCA	GACAAGGATG	GAGACCTCAT	600
٠	TGCCACCAAG	GAGGAGTTCA	CAGCTTTCCT	GCACCCTGAG	GAGTATGACT	ACATGAAAGA	660
20	TATAGTAGTA	CAGGAAACAA	TGGAAGATAT	AGATAAGAAT	GCTGATGGTT	TCATTGATCT	720
	AGAAGAGTAT	ATTGGTGACA	TGTACAGCCA	TGATGGGAAT	ACTGATGAGC	CAGAATGGGT	780
	AAAGACAGAG	CGAGAGCAGT	TTGTTGAGTT	TCGGGATAAG	AACCGTGATG	GGAAGATGGA	840
25	CAAGGAAGAG	ACCAAAGACT	GGATCCTTCC	CTCAGACTAT	GATCATGCAG	AGGCAGAAGC	900
	CAGGCACCTG	GTCTATGAAT	CAGACCAAAA	CAAGGATGGC	AAGCTTACCA	AGGAGGAGAT	960
30	CGTTGACAAG	TATGACTTAT	TTGTTGGCAG	CCAGGCCACA	GATTTTGGGG	AGGCCTTAGT	1020
	ACGGCATGAT	GAGTTCTGAG	CTRCGGAGGA	ACCCTCATTT	CCTCAAAAGT	AATTTATTTT	1080
	TACAGCTTCT	GGTTTCACAT	GAAATTGTTT	GCGCTACTGA	GACTGTTACT	ACAAACTTTT	1140
35	TAAGACATGA	AAAGGCGTAA	TGAAAACCAT	CCCGTCCCCA	TTCCTCCTCC	TCTCTGAGGG	1200
	ACTGGAGGGA	AGCCGTGCTT	CTGAGGAACA	ACTCTAATTA	GTACACTTGT	GTTTGTAGAT	1260
40	TTACACTTTG	TATTATGTAT	TAACATGGCG	TGTTTATTTT	TGTATTTTTC	TCTGGTTGGG	1320
	AGTATGATAT	GAAGGATCAA	GATCCTCAAC	TCACACATGT	AGACAAACAT	TAGCTCTTTA	1380
	CTCTTTCTCA	ACCCCTTTTA	TGATTTTAAT	AATTCTCACT	TAACTAATTT	TGTAAGCCTG	1440
45	AGATCAATAA	GAAATGTTCA	GGAGAGAGGA	AAGAAAAAA	ATATATGCTC	CACAATTTAT	1500
	ATTTAGAGAG	AGAACACTTA	GTCTTGCCTG	TCAAAAAGTC	CAACATTTCA	TAGGTAGTAG	1560
50	GGGCCACATA	TTACATTCAG	TTGCTATAGG	TCCAGCAACT	GAACCTGCCA	TTACCTGGGC	1620
	AAGGAAAGAT	CCCTTTGCTC	TAGGAAAGCT	TGGCCCAAAT	TGATTTTCTT	CTTTTTCCCC	1680
	CTGTAGGACT	GACTGTTGGC	TAATTTTGTC	AAGCACAGCT	GTGGTGGGAA	GAGTTAGGGC	1740
55	CAGTGTCTTG	AAAATCAATC	AAGTAGTGAA	TGTGATCTCT	TTGCAGAGCT	ATAGATAGAA	1800
	ACAGCTGGAA	AACTAAAGGA	AAAATACAAG	TGTTTTCGGG	GCATACATTT	TTTTTCTGGG	1860
60	TGTGCATCTG	TIGAAATGCT	CAAGACTTAA	TTATTTGCCT	TTTGAAATCA	CTGTAAATGC	1920

	CCCCATCCGG	TTCCTCTTCT	TCCCAGGTGT	GCCAAGGAAT	TAATCTTGGT	TTCACTACAA	1980
	ттааааттса	CTCCTTTCCA	ATCATGTCAT	TGAAAGTGCC	TTTAACGAAA	GAAATGGTCA	2040
5	CTGAATGGGA	ATTCTCTTAA	GAAACCCTGA	GATTAAAAAA	AGACTATTTG	GATAACTTAT	2100
	AGGAAAGCCT	AGAACCTCCC	AGTAGAGTGG	GGATTTTTTT	CTTCTTCCCT	TTCTCTTTTG	2160
10	GACAATAGTT	AAATTAGCAG	TATTAGTTAT	GAGTTTGGTT	GCAGTGTTCT	TATCTTGTGG	2220
10	GCTGATTTCC	AAAAACCACA	TGCTGCTGAA	TTTACCAGGG	ATCCTCATAC	CTCACAATGC	2280
	AAACCACTTA	CTACCAGGCC	TTTTTCTGTG	TCCACTGGAG	AGCTTGAGCT	CACACTCAAA	2340
15	GATCAGAGGA	CCTACAGAGA	GGGCTCTTTG	GTTTGAGGAC	CATGGCTTAC	CTTTCCTGCC	2400
	TTTGACCCAT	CACACCCCAT	TTCCTCCTCT	TTCCCTCTCC	CCGCTGCCAA	TTCCTGCAGC	2460
20	CCGGGGGAAC	CACTAGTT					2478
20							

(2) INFORMATION FOR SEQ ID NO: 91:

25

30

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2058 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 91:

TCGGCCTTGC TTTTGTGGYC TTCCTCTGTG GCCAGAGCGT TTTCATCACC AAGCCTCCTG 60 35 ATGGCAGTNC CTTCACCGAT ATGTTCAAGA TACTGACGTA TTCCTGCTGT TCCCAGAAGC 120 GAAGTGGAGA GCGCCAGAGT AATGGTGAAG GCATTGGAGT NTTTCAGCAA TCTTCTAAAC 180 240 AAAGTCTGTT TGATTCATGT AAGATGTCTC ATGGTGGGCC ATTTACAGAA GAGAAAGTGG 40 AAGATGTGAA AGCTCTGGTC AAGATTGTCC CTGTTTTCTT GGCTTTGATA CCTTACTGGA 300 CAGTGTATTT CCAAATGCAG ACAACATATG TTTTACAGAG TCTTCATTTG AGGATTCCAG 360 45 AAATTTCAAA TATTACAACC ACTCCTCACA CGCTCCCTGC AGCCTGGCTG ACCATGTTTG 420 ATGCTGTGCT CATCCTCCTG CTCATCCCTC TGAAGGACAA ACTGGTCGAT CCCATTTTGA 480 GAAGACATGG CCTGCTCCCA TCCTCCCTGA AGAGGATCGC CGTGGGCATG TTCTTTGTCA 540 50 TGTGCTCRGC CTTTGCTGCA GGAATTTTGG AGAGTAAAAG GCTGAACCTT GTTAAAGAGA 600 AAACCATTAA TCAGACCATC GGCAACGTCG TCTACCATGC TGCCGATCTG TCGCTGTGGT 660 55 GGCAGGTGCC GCAGTACTTG CTGATTGGGA TCAGCGAGAT CTTTGCAAGT ATCGCAGGCC 720 TGGAATTTGC ATACTCAGCT GCCCCCAAGT CCATGCAGAG TGCCATAATG GGCTTGTTCT 780 TTTTCTTCTC TGGCGTCGGG TCGTTCGTGG GTTCTGGACT GCTGGCACTG GTGTCTATCA 840 60

	ANGCCATCGG	AIGGAIGAGC	AGICACACAG	ACTITIGGIAA	TAT TAACGGC	TGCTATTIGA	900
5	ACTATTACTT	TTTCCTTCTG	GCTGCTATTC	AAGGAGCTAC	CCTCCTGCTT	TTCCTCATTA	960
	TTTCTGTGAA	ATATGACCAT	CATCGAGACC	ATCAGCGATC	AAGAGCCAAT	GGCGTGCCCA	1020
	CCAGCAGGAG	GGCCTGACCT	TCCTGAGGCC	ATGTGCGGTT	TCTGAGGCTG	ACATGTCAGT	1080
10	AACTGACTGG	GGTGCACTGA	GAACAGGCAA	GACTITAAAT	TCCCATAAAA	TGTCTGACTT	1140
	CACTGAAACT	TGCATGTTGC	CTGGATTGAT	TTCTTCTTTC	CCTCTATCCA	AAGGAGCTTG	1200
15	GTAAGTGCCT	TACTGCAGCG	TGTCTCCTGG	CACGCTGGGC	CCTCCGGGAG	GAGAGCTGCA	1260
	GATTTCGAGT	ATGTCGCTTG	TCATTCAAGG	TCTCTGTGAA	TCCTCTAGCT	GGGTTCCCTT	1320
	TTTTACAGAA	ACTCACAAAT	GGAGATTGCA	AAGTCTTGGG	GAACTCCACG	TGTTAGTTGG	1380
20	CATCCCAGTT	TCTTAAACAA	ATAGTATCAC	CTGCTTCCCA	TAGCCATATC	TCACTGTAAA	1440
	AAAAAAATT	AATAAACTGT	TACTTATATT	TAAGAAAGTG	AGGATTTTTT	TTTTTTAAAG	1500
25	ATAAAAGCAT	GGTCAGATGC	TGCAAGGATT	TTACATAAAT	GCCATATTTA	TGGTTTCCTT	1560
	CCTGAGAACA	ATCTTGCTCT	TGCCATGTTC	TTTGATTTAG	GCTGGTAGTA	AACACATTTC	1620
	ATCTGCTGCT	TCAAAAAGTA	CTTACTTTT	AAACCATCAA	CATTACTTTT	CTTTCTTAAG	1680
30	GCAAGGCATG	CATAAGAGTC	ATTTGAGACC	ATGTGTCCCA	TCTCAAGCCA	CAGAGCAACT	1740
	CACGGGGTAC	TTCACACCTT	ACCTAGTCAG	AGTGCTTATA	TATAGCTTTA	TTTTGGTACG	1800
35	ATTGAGACTA	AAGACTGATC	ATGGTTGTAT	GTAAGGAAAA	CATTCTTTTG	AACAGAAATA	1860
	GTGTAATTAA	AAATAATTGA	AAGTGTTAAA	TGTGAACTTG	AGCTGTTTGA	CCAGTCACAT	1920
	TTTTGTATTG	TTACTGTACG	TGTATCTGGG	GCTTCTCCGT	TTGTTAATAC	TTTTTCTGTA	1980
40	TTTGTTGCTG	TATTTTTGGC	ATAACTTTAT	TATAAAAAGC	ATCTCAAATG	CGAAAWAAAA	2040
	АААААААА	AAAAAAAC					2058
45							
	(2) INFORMA	ATION FOR SE	Q ID NO: 92	l:			
	(i)	SEQUENCE CH	HARACTERIST	ICS:			
50			GTH: 1411 b E: nucleic a				
		(C) STR	ANDEDNESS: O	double			
55	(xi)) SEQUENCE I	DESCRIPTION	: SEQ ID NO:	: 92:		
	GGCACAGGAG	CGACCCGGGA	GAAGGAGGGC	CAMGAKGCGG	AAGCGGAGGA	GTCTCCAGGA	60
60					TTTCAGATAT		120
50							

	CAAGACATCC CCGTAGAAGG AGAAATCACC ATTCCTATGA GATCTCGCAT CCGGGAGTTT	180
	GACAGCTCCA CATTAAATGA ATCTGTTCGC AATACCATCA TGCGTGATCT AAAAGCTGTT	240
5	GGGAAAAAAT TCATGCATGT TTTGTACCCA AGGAAAAGTA ATACTCTTTT GAGAGATTGG	300
	GATTTGTGGG GCCCTTTGAT CCTTTGTGTG ACACTCGCAT TAATGCTGCA AAGAGACTCT	360
	GCAGATAGTG AAAAAGATGG AGGGCCCCAA TTTGCAGAGG TGTTTGTCAT TGTCTGGTTT	420
10	GGTGCAGTTA CCATCACCCT CAACTCAAAA CTTCTTGGAG GGAACATATC TTTTTTTCAG	480
	AGCCTCTGTG TGCTGGGTTA CTGTATACTT CCCTTGACAG TAGCAATGCT GATTTGCCGG	540
15	CTGGTACTTT TGGCTGATCC AGGACCTGTA AACTTCATGG TTCGGCTTTT TGTGGTGATT	600
	GTGATGTTTG CCTGGTCTAT AGTTGCCTCC ACAGCTTTCC TTGCTGATAG CCAGCCTCCA	660
20	AACCGCAGAG CCCTAGCTGT TTATCCTGTT TTCCTGTTTT ACTTTGTCAT CAGTTGGATG	720
20	ATTCTCACCT TTACTCCTCA GTAAATCAGG AATGGGAAAT TAAAAAACCAG TGAATTGAAA	780
	GCACATCTGA AAGATGCAAT TCACCATGGA GCTTTGTCTC TGGCCCTTAT TTGTCTAATT	840
25	TTGGAGGTAT TTGATAACTG AGTAGGTGAG GAGATTAAAA GGGAGCCATA TAGCACTGTC	900
	ACCCCTTATT TGAGGAACTG ATGTTTGAAA GGCTGTTCTT TTCTCTCTTA ATGTCATTTC	960
20	TTTAAAAATA CATGTGCATA CTACACACAG TATATAATGC CTCCTTAAGG CATGATGGAG	1020
30	TCACCGTGGT CCATTTGGGT GACAACCAGT GACTTGGGAA GCACATAGAT ACATCTTACA	1080
	AGTTGAATAG AGTTGATAAC TATTTTCAGT TTTGAGAATA CCAGTTCAGG TGCAGCTCTT	1140
35	AAACACATTG CCTTATGACT ATTAGAATAT GCCTCTCTTT TCATAAATAA AAATACATGG	1200
	TCTATATCCA TTTTCTTTTA TTTCTCTCTC TTAAGCTTAA AAAGGCAATG AGAGAGGTTA	1260
40	GGAGTGGGTT CATACACGGA GAATGAGAAA ACATGCATTA ACCAATATTC AGATTTTGAT	1320
40	CAGGGGAAAT TCTAYACTTG TTGCAAAAAA AAAAAAAAAA AAACTCGAGG GGGGCCCGGT	1380
	ACCCAATCGC NGTATATGAT CGNAAACAAT C	1411
45		
	(2) INFORMATION FOR SEQ ID NO: 93:	
5 0		
50	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 2187 base pairs	
	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
	(D) TOPOLOGY: linear	
55	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 93:	
	GCTTTGGCTT TTTTTGGCGG ACTGGGGCGC CCTCCGGAAG CGTTTCCAAC TTTCCAGAAG	60
60	THE THE PROPERTY OF THE PROPERTY OF THE PROPERTY THE ATTOCKED AND THE PROPERTY OF THE PROPERTY	120

	GCGGGCTAAC	AGTAGAATCG	TGTCGCGCTC	GAGAGCGAGA	GTCACGTCCC	GGCGCTAGCC	180
5	CAGCCCGACC	: CAGGCCCACC	GTGGTGCACC	CAAACCACTI	CCTGGCCATG	CGCTCCCTCC	240
	TGCTTCTCAG	CGCCTTCTGC	CTCCTGGAGG	CGGCCCTGGC	CGCCGAGGTG	AAGAAACCTG	300
	CAGCCGCAGC	AGCTCCTGGC	ACTGCGGAGA	AGTTGAGCCC	CAAGGCGGCC	ACGCTTGCCG	360
10	AGCGCAGCCG	GCCTGGCCTT	CAGCTTGTAC	CAGGCCATGG	CCAAGGACCA	GGCAGTGGAG	420
	AACATCCTGG	TGTCACCCGT	GGTGGTGGCC	TCGTCGCTGG	GGCTCGTGTC	CCTGGGCGGC	480
15	AAGGCGACCA	CGGCGTCGCA	GGCCAAGGCA	GTGCTGAGCG	CCGAGCAGCT	GCGCGACGAG	540
	GAGGTGCACG	CCGGCCTGGG	CGAGCTGCTG	CGCTCACTCA	GCAACTCCAC	GGCGCGCAAC	600
	GTGACCTGGA	AGCTGGGCAG	CCGACTGTAC	GGACCCAGCT	CAGTGAGCTT	CGCTGATGAC	660
20	TTCGTGCGCA	GCAGCAAGCA	GCACTACAAC	TGCGAGCACT	CCAAGATCAA	CTTCCGCGAC	720
	AAGCGCAGCG	CGCTGCAGTC	CATCAACGAG	TGGGCCGCGC	AGACCACCGA	CGGCAAGCTG	780
25	CCCGAGGTCA	CCAAGGACGT	GGAGCGCACG	GACGGCGCCC	TGTTAGTCAA	CGCCATGTTC	840
	TTCAAGCCAC	ACTGGGATGA	GAAATTCCAC	CACAAGATGG	TGGACAACCG	TGGCTTCATG	900
	GTGACTCGGT	CCTATACCGT	GGGTGTCATG	ATGATGCACC	GGACAGGCCT	CTACAACTAC	960
30	TACGACGACG	AGAAGGAAAA	GCTGCAAATC	GTGGAGATGC	CCCTGGCCCA	CAAGCTCTCC	1020
	AGCCTCATCA	TCCTCATGCC	CCATCACGTG	GAGCCTCTCG	AGCGCCTTGA	AAAGCTGCTA	1080
35	ACCAAAGAGC	AGCTGAAGAT	CTGGATGGGG	AAGATGCAGA	AGAAGGCTGT	TGCCATCTCC	1140
	TTGCCCAAGG	GTGTGGTGGA	GGTGACCCAT	GACCTGCAGA	AACACCTGGC	TGGGCTGGGC	1200
	CTGACTGAGG	CCATTGACAA	GAACAAGGCC	GACTTGTCAC	GCATGTCAGG	CAAGAAGGAC	1260
40	CTGTACCTGG	CCAGCGTGTT	CCACGCCACC	GCCTTTGAGT	TGGACACAGA	TGGCAACCCT	1320
	TTGACCAGAA	TTACGGGCGG	AGGAGTGCGC	ACCCAAGTGT	TCTACGCCGA	CCACCCCTTC	1380
45	ATTTCCTAGT	GCGGGACACC	CAAAGCGGTC	CCTGCTATTC	ATTGGGCGCC	TGGTCCGGCC	1440
	TAAGGGTGAC	AAGATGCGAG	ACGAGTTATA	GGCCTCAGGG	TGCACACAGG	ATGGCAGGAG	1500
	GCATCCAAAG	GCTCCTGAGA	CACATGGGTG	CTATTGGGGT	TGGGGGGAG	GTGAGGTACC	1560
50	AGCCTTGGAT	ACTCCATGGG	GTGGGGTGGA	AAAGCAGACC	GGGGTTCCCG	TGTGCCTGAG	1620
	CGGACTTCCC	AGCTAGAATT	CACTCCACTT	GGACATGGGC	CCCAGATACC	ATGATGCTGA	1680
55	GCCCGGAAAC	TCCACATCCT	GTGGGACCTG	GGCCATAGTC	ATTCTGCCTG	CCCTGAAAGT	1740
	CCCAGATCAA	GCCTGCCTCA	ATCAGTATTC	ATATTTATAG	CCAGGTACCT	TCTCACCTGT	1800
	GAGACCAAAT	TGAGCTAGGG	GGGTCAGCCA	GCCCTCTTCT	GACACTAAAA	CACCTCAGCT	1860
60	GCCTCCCCAG	CTCTATCCCA .	ACCTCTCCCA	ACTATAAAAC	TAGGTGCTGC	AGCCCCTGGG	1920

	ACCAGGCACC CCCAGAATGA CCTGGCCGCA GTGAGGCGGA TTGAGAAGGA GCTCCCAGGA	1980
_	GGGGCTTCTG GGCAGACTCT GGTCAAGAAG CATCGTGTCT GGCGTTGTGG GGATGAACTT	2040
5	TTTGTTTTGT TTCTTCCTTT TTTAGTTCTT CAAAGATAGG GAGGGAAGGG GGAACATGAG	2100
	CCTTTGTTGC TATCAATCCA AGAACTTATT TGTACATTTT TTTTTTCAAT AAAACTTTTC	2160
10	CAATGACAAA AAAAAAAAA AAAAAAA	2187
15	(2) INFORMATION FOR SEQ ID NO: 94:	
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 757 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double	
20	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 94:	
25	GACAGTACGG TCGGATTCCC GGGTCGACCC ACGCGTCCGC GGACGGTGAA GAAGGTGAAG	60
	ATGGCGGTGG CCAGGGCCGG GGTCTTGGGA GTCCAGTGGC TGCAAAGGGC ATCCCGGAAC	120
30	GTGATGCCGC TGGGCGCACG GACAGCCTCC CACATGACCA AGGACATGTT CCCGGGGCCC	180
30	TATCCTAGGA CCCCAGAAGA ACGGGCCGCC GCCGCCAAGA AGTATAATAT GCGTGTGGAA	240
	GACTACGAAC CTTACCCGGA TGATGGCATG GGGTATGGCG ACTACCCGAA GCTCCCTGAC	300
35	CGCTCACAGC ATGAGAGAGA TCCATGGTAT AGCTGGGACC AGCCGGGCCT GAGGTTGAAC	360
	TGGGGTGAAC CGATGCACTG GCACCTAGAC ATGTACAACA GGAACCGTGT GGATACATCC	420
40	CCCACACCTG TTTCTTGGCA TGTCATGTGT ATGCAGCTCT TCGGTTTCCT GGCTTTCATG	480
	ATATTCATGT GCTGGGTGGG GGACGTGTAC CCTGTCTACC AGCCTGTGGG ACCAAAGCAG	540
	TATCCTTACA ATAATCTGTA CCTGGAACGA GGCGGTGATC CCTCCAAAGA ACCAGAGCGG	600
45	GTGGTTCACT ATGAGATCTG AGGAGGCTTC GTGGGCTTTT GGGTCCTCTA ACTAGGACTC	660
	CCTCATTCCT AGAAATTTAA CCTTAATGAA ATCCCTAATA AAACTCAGTG CTGTGTTAAA	720
50	AAAAAAAA AAAAAAAA AAAAAGGGGG GCCCCNN	757
55	(2) INFORMATION FOR SEQ ID NO: 95:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 2394 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double	
60	(D) TOPOLOGY: linear	

(xi)	SEQUENCE	DESCRIPTION:	SEQ	ID NO:	95:

5	GGCACGAGCA	CTCCTGCACT	TCCCCACCCC	CACGACCGAA	CCTGGCTTCG	CTAACGCCCT	60
J	CCCAGCTCCC	TCGGGCCTGA	CTTCCGGTTT	CCTCGCGCGT	CCCTGGCGCC	GAGCCGCGGA	120
	CAGCAGCCCC	TTTTCCGGCT	GAGAGCTCAT	CCACACTTCC	AATCACTTTC	CGGAGTGCTT	180
10	CCCCTCCCTC	CGGCCCGTGC	TGGTCCCGAC	GGCGGGCCTG	GGTCTCGCGC	GCGTATTGCT	240
	GGGTAACGGG	CCTTCTCYCG	CGTCGGCCCG	GCCCCTCCTG	CCTCGGCTCG	TCCCTCCTTC	300
15	CAGAACGTCC	CGGGCTCCTG	CCGAGTCAGA	AGAAATGGGA	CTCCCTCCGC	GACGTGCCCG	360
	GAGCAGCTCC	CTTCGCTGTG	GAAGCGGCGG	TGTCTTCGAA	GAAACCGGAA	GCCCGTGGTG	420
	ACCCCTGGCG	ACCCGGTTTG	TTTTCGGTCC	GTTTCCAAAC	ACTAAGGAAT	CGAAACTCGG	480
20	CGGCCTTGGG	GGCGGCCCTA	CGTAGCCTGG	CTTCTGGTTG	TCATGGATGC	ACTGGTAGAA	540
	GATGATATCT	GTATTCTGAA	TCATGAAAAA	GCCCATAAGA	GAGATACAGT	GACTCCAGTT	600
25	TCAATATATT	CAGGAGATGA	ATCTGTTGCT	TCCCATTTTG	CTCTTGTCAC	TGCATATGAA	660
	GACATCAAAA	AACGACTTAA	GGATTCAGAG	AAAGAGAACT	CTTTGTTAAA	GAAGAGAATA	720
	AGATTTTTGG	AAGAAAAGCT	AATAGCTCGA	TTTGAAGAAG	AAACAAGTTC	CGTGGGACGA	780
30	GAACAAGTAA	ATAAGGCCTA	TCATGCATAT	CGAGAGGTTT	GCATTGATAG	AGATAATTTG	840
	AAGAGCAAAC	TGGACAAAAT	GAATAAAGAC	AACTCTGAAT	CTTTGAAAGT	ATTGAATGAG	900
35	CAGCTACAAT	CTAAAGAAGT	AGAACTCCTC	CAGCTGAGGA	CAGAGGTGGA	AACTCAGCAG	960
	GTGATGAGGA	ATTTAAATCC	ACCTTCATCA	AACTGGGAGG	TGGAAAAGTT	GAGCTGTGAC	1020
	CTGAAGATCC	ATGGTTTGGA	ACAAGAGCTG	GAACTGATGA	GGAAAGAATG	TAGCGATCTC	1080
40	AAAATAGAAC	TACAGAAAGC	CAAACAAACG	GATCCATATC	AGGAAGACAA	TCTGAAGAGC	1140
	AGAGATCTCC	AAAAACTAAG	CATTTCAAGT	GATAATATGC	AGCATGCATA	CTGGGAACTG	1200
45	AAGAGAGAAA	TGTCTAATTT	ACATCTGGTG	ACTCAAGTAC	AAGCTGAACT	ACTAAGAAAA	1260
	CTGAAAACCT	CAACTGCAAT	CAAGAAAGCC	TGTGCCCCTG	TAGGATGCAG	TGAAGACCTT	1320
	GGAAGAGACA	GCACAAAACT	GCACTTGATG	AATTTTACTG	CAACATACAC	AAGACATCCC	1380
50	CCTCTCTTAC	CAAATGGCAA	AGCTCTTTGT	CATACCACAT	CTTCCCCTTT	ACCAGGAGAT	1440
	GTAAAGGTTT	TATCAGAGAA	AGCAATCCTC	CAATCATGGA	CAGACAATGA	GAGATCCATT	1500
55	CCTAATGATG	GTACATGCTT	TCAGGAACAC	AGTTCTTATG	GCAGAAATTC	TCTGGAAGAC	1560
	AATTCCTGGG	TATTTCCAAG	TCCTCCTAAA	TCAAGTGAGA	CAGCATTTGG	GGAAACTAAA	1620
	ACTAAAACTT	TGCCTTTACC	CAACCTTCCA	CCACTGCATT	ACTTGGATCA	ACATAATCAG	1680
60	AACTGCCTTT	ATAAGAATTA	ATTTGGAAGA	GATTCACGAT	TTCACCATGA	GGACACTTAT	1740

	CTCTTTCAGT GGTCCTCCCA AGAAATTATT TAACAAACTG AANGGAGATT TTGATTAAAA	1800
_	TTTTGCAGAG GTCTTCAGTA TCTATATTTG AACACACTGT ACAATAGTAC AAAAACCAAC	1860
5	ATAGTTGGTT TTCTAGTATG AAAGAGCACC CTCTAGCTCC ATATTCTAAG AATCTGAAAT	1920
	ATGCTACTAT ACTAATTAAT AAGTAAACTT AAGGTGTTTA AAAAACTCTG CCTTCTATAT	1980
10	TAATTGTAAA ATTTTGCCTC TCAGAAGAAT GGAATTGCAG ATTGTAGACG TGGTTTTACA	2040
	AAATGTGAAA TGTCTAAATA TCTGTTCATA AAAATAAAAG GAAAACATGT TTCTTCAAAT	2100
	TGCATAATGG AACAAATGGC AATGTGAGTA GGTTACATTT CTGTTGTTAT AATGCGTAAA	2160
15	GATATTGAAA ATATAATGAA ATAAAAGCAT CTTAGGTTAT ACCATCTTTA TATGCTATTG	2220
	CGTTICAATA TTTAAGATTT AAAGTGATTT TTTGGTCACA GTGTTTTGTT GATAAAATTT	2280
20	TTTTAGAATT GAAGTTTGAA TTCTAAGACT TGAAACAACC TGATCACTGA AGCCAACTTT	2340
	GTCCCAGCAC ATTCCTTAAG TCCTAATTGG GGAAAAAAAA AAAAAAAAAC TCGA	2394
25		
25		
	(2) INFORMATION FOR SEQ ID NO: 96:	
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 672 base pairs	
50	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NC: 96:	
	AGTGCTCTGT TGCCCAGGCT GGAGTGCGTT AGTGTAATGT CAGTCCACTG CAACCTCCAC	60
	CCCCAGGTTC AAGCAATTCT CATGCCTCAG CCTCCCAAGT AGCTGAAATT ACTGGCATGC	120
40	ACCACCACAC CCAGCTGATG TTTATTTATT TATTTATATA TTTATTTATT TTAGGTGTTT	180
	TYPPPPPTT TYPPGAGAC GGAGTCTTGC TCTGTTGCCC TGGGTGTGGT TACGTGGRAT	240
45	TACCATYCTG GGTGACTCAC TGAAATGTAC TCMCAGTGAG TCATGCCTTC MAATGACATC	300
	TCAAGTTCTG CCTGCTTGGA GATACATCTG GGGATCTTAA GGGGTGAGGG ACTACTCAAC	360
	AAGAAGGAAT TTAGCCTGTC TTTTTAAATA AACGGCATTT CTTTTTCCTA KAAAAATGGG	420
50	AAATTCTTCA ATTCTCTAAT ACAGGGACAC TGAGATAACA AAGAGGAAAG TGTCTGGTTG	480
	GAGGTTGGGA RGCCACCCTG GGGTCTCTCC TACAAAAATG GAAAAGAAAA	540
55	AAATCMAGCA AAGCACAARA AAKTTTCCCT TTGCTAAAAG GGAAAAGATG CCCCMCAATG	600

CCCATAAACA TGAACTGGGG ATAAGGAGGA RAATGTCTCT YCTTGGCACC CCCAAACAAA

60

CGTTAATTAC CC

1080

1140

1200

1260

1320

1380

1419

5	(2) INFORMA	TION FOR SI	EQ ID NO: 9	7:			
10	(i) :	(A) LEN (B) TYP (C) STR	HARACTERIST GTH: 1419 b E: nucleic ANDEDNESS: OLOGY: line	ase pairs acid double			
	(xi)	SEQUENCE I	DESCRIPTION	: SEQ ID NO	: 97:		
15	TAAGAACAGA A	ACAGCAAGTA	TGAACCACAT	GGAACTTAAA	ACATATGGGT	GTGAAGTCCA	60
	CTTATGTAGA (CAAAACTTAT	AATTTCCAAA	CTGTTGTCTA	GTATACAGTG	ATCAGTTGCT	120
	CTCTGTTCAA C	STCATTCCAC	ACATTTCCCT	ATTTTAGGCT	ATTATAATAT	AGAAAGAAAA	180
20	TGGGAAGCAT I	TAGTTGGAGC	TAGAAAATGA	ACTGTATATT	ATTGCTATAT	TTGCTAATAC	240
	CAACTATTTC A	ATAAGTGTT	GTACCATATG	TAGCATTAAA	TATAAAATAC	ATAAAAGAAT	300
25	GTACAGAAAA T	AGCTTTTAT	TGAGTAATAT	TACATTTCAT	TTATACTGTA	GCAATATATT	360
	TGTAGGTATA C	TCTGTAAGG	GCTTTAAATA	AAAGAGGTCC	ATTAATACTT	CCTTATAAAA	420
	ATTCTAGTCT G	STTTCATTAC	TGCCCAGATG	TTTTAGAGAT	AAATATTTAT	GCAGAAGGTA	480
30	TTTTKGAAAG T	CYCCYTTTG	TCTGATAGAG	TTTAACNAGA	TATTTAAATT	TAGTGCYCNA	540
	GAAATCCCAC A	AGTCACGGT	CTAAACACAC	TTAGAATACT	ACAGCATAAA	TCTGTTAGCA	600
35	TTANTTGCCA A	ATAAGACAG	TTGGGATCCC	AAACCCCAAG	TCCTTGAGCA	ATGTTTTCC	660
	TCAAAAAGCT G	CTATNCCAA	TGATATAGGA	AAAWACATTG	TGTTTTCCTA	AACACACTTT	720
	TCTTTTTAAA T	GTGCTTCAT	TGTTTGATTT	GGTCCTGCCT	AAATTTCACA	AGCTAGGCCA	780
40	ATGAAGGCTG A	ATCAAAGAC	ATTTCATCCA	CCAATATCAT	GTGTAGATAT	TATGTATAGA	840
	AAATAAAATA A	ATTATGGCT	CTAACTTCTG	TGTTGCTGTT	TATCTTGTTA	TTTTTCGGCG	900
45	TTATACTAAT G						960
	GTCTGWATTT T	GSTCCTTAG	ATGKGAATAT	TTCTTATTAG	TYTGCTYCCT	GCWACGCAAT	1020

GACTGCATTT CTATCATTTC TCAGTTTGTT AGWATATGTG GATAGTATTC TACTGTATAA

ATGATTGCAA AGTTTATCAA AAACAAATTA TTATATGTAG CTTTTCTACA GTGCTTTGCT

AAACCATGTA GTACTAGTTA AGTSTTCCTT GAAAATAAAG ATACACTCTT ATAGGGGACA

GTTCCTGTTC ACTCCCAGGA AACTTTTTTA AAAGATGACA CTGAATGTTT ATTGCACTTT

AGTGCAGTGA AGTGGCAATA AAACCTAACA TGAATCAAGG TTGTTTATGG CAGATGCATG

TGTTGCTTTA CAGAGTTTAG CAAAAGCTCT TAATTTTATG TCATACTGTA TTCTACTGAA

TAATAAAGCT AACATTATTC AATAATAAAA TGGAAAAAA

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5	(2)	INFORMATION	FOR	SEO	ID	NO:	98:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1487 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 98:

	, , <u>-</u> <u>-</u> <u>-</u>	
15	GCGACCGCGC CCCTTTCAGC TAGCTCGCTC GCTCGCTCTG CTTCCCTGCT GCCGGCTGCG	60
	CATGGCKWTG GCGTTGGCGG CGCTGGCGGC GGTCGAGCCG GCCTGCGCAG CCGGTACCAG	120
20 25	CAGTTGCAGA ATGAAGAAGA GTCTGGAGAA CCTGAACAGG CTGCAGGTGA TGCTCCTCCA	180
	CCTTACAGCA GCATTTCTGC AGAGAGCGCA GTTTTCCACC TATTTCCCTG GATATTTTGA	240
	TGGTCAGTAC TGGCTCTGGT GGGTGTTCCT TGTTTTAGGC TTTCTCCTGT TTCTCAGAGG	300
	ATTTATCAAT TATGCAAAAG TTCGGAAGAT GCCAGAAACT TTCTCAAATC TCCCCAGGAC	360
	CAGAGTTCTC TTTATTTATT AAAGATGTTT TCTGGCAAAG GCCTTCCTGC ATTTATGAAT	420
30	TCTCTCTCAA GAAGCAAGAG AACACCTGCA GGAAGTGAAT CAAGATGCAG AACACAGAGG	480
	AATAATCACC TGCTTTAAAA AAATAAAGTA CTGTTGAAAA GATCATTTCT CTCTATTTGT	540
	TCCTAGGTGT AAAATTTTAA TAGTTAATGC AGAATTCTGT AATCATTGAA TCATTAGTGG	600
35	TTAATGTTTG AAAAAGCTCT TGCAATCAAG TCTGTGATGT ATTAATAATG CCTTATATAT	660
	TGTTTGTAGT CATTTTAAGT AGCATGAGCC ATGTCCCTGT AGTCGGTAGG GGGCAGTCTT	720
40	GCTTTATTCA TCCTCCATCT CAAAATGAAC TTGGAATTAA ATATTGTAAG ATATGTATAA	780
	TGCTGGCCAT TTTAAAGGGG TTTTCTCAAA AGTTAAACTT TTGTTATGAC TGTGTTTTTG	840
45	CACATAATCC ATATTTGCTG TTCAAGTTAA TCTAGAAATT TATTCAATTC TGTATGAACA	900
	CCTGGAAGCA AAATCATAGT GCAAAAATAC ATTTAAGGTG TGGTCAAAAA TAAGTCTTTA	960
	ATTGGTAAAT AATAAGCATT AATTTTTAT AGCCTGTATT CACAATTCTG CGGTACCTTA	1020
50	TTGTACCTAA GGGATTCTAA AGGTGTTGTC ACTGTATAAA ACAGAAAGCA CTAGGATACA	1080
	AATGAAGCTT AATTACTAAA ATGTAATTCT TGACACTCTT TCTATAATTA GCGTTCTTCA	1140
55	CCCCCACCC CACCCCCACC CCCCTTATTT TCCTTTTGTC TCCTGGTGAT TAGGCCAAAG	1200
	TCTGGGAGTA AGGAGGGAT TAGGTACTTA GGAGCAAAGA AAGAAGTAGC TTGGAACTTT	1260
	TGAGATGATC CCTAACATAC TGTACTACTT GCTTTTACAA TGTGTTAGCA GAAACCAGTC	1320
	GGTTATAATG TAGAATGATG TGCTTTCTGC CCAAGTGGTA ATTCATCTTG GTTTGCTATG	1380
60		

	TTAAAACTGT AAATACAACA GAACATTAAT AAATATCTCT TGTGTAGCAC CTTTAAAAAA	1440
	AAAAAAAAA AAAAAAAAAAAAAAAAAAAAAAAAAAAA	1487
5		
	(2) INFORMATION FOR SEQ ID NO: 99:	
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 1653 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double	
15	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 99:	
	GCGACCGCGC CCTTCAGCTA GCTCGCTCGC TCGCTCTGCT TCCCTGCTGC CGGCTGCGCA	60
20	TGGCTTNGGC GTTGGCGGCG CTGGCGGCGG CTCGAGCCGC CTGCGSAGCC GGTACCAGCA	120
	GTTGCAGAAT GAAGAAGAGT CTGGAGAACC TGAACAGGCT GCAGGTGATG CTCCTCCACC	180
25	TTACAGCAGC ATTTCTGCAG AGAGCGCACA TNATTTTGAC TACAAGGATG AGTCTGGGTT	240
	TCCAAAGCCC CCATCTTACA ATGTAGCTAC AACACTGCCC AGTTATGATG AAGCGGAGAG	300
	GACCAAGGCT GAAGCTACTA TCCCTTTGGT TCCTGGGAGA GATGAGGATT TTGTGGGTCG	360
30	GGATGATTTT GATGATGCTG ACCAGCTGAG GATAGGAAAT GATGGGATTT TCATGTTAAC	420
	TTTTTTCATG GCATTCCTCT TTAACTGGAT TGGGTTTTTC CTGTCTTTTT GCCTGACCAC	480
35	TTCAGCTGCA GGAAGGTATG GGGCCATTTC AGGATTTGGT CTCTCTCTAA TTAAATGGAT	540
33	CCTGATTGTC AGGTTTTCCA CCTATTTCCC TGCATTTATG AATTCTCTCT CAAGAAGCAA	600
	GAGAACACCT GCAGGAAGTG AATCAAGATG CAGAACACAG AGGAATAATC ACCTGCTTTA	660
40	AAAAAATAAA GTACTGTTGA AAAGATCATT TCTCTCTATT TGTTCCTAGG TGTAAAATTT	720
	TAATAGTTAA TGCAGAATTC TGTAATCATT GAATCATTAG TGGTTAATGT TTGAAAAAGC	780
15	TCTTGCAATC AAGTCTGTGA TGTATTAATA ATGCCTTATA TATTGTTTGT AGTCATTTTA	840
45	AGTAGCATGA GCCATGTCCC TGTAGTCGGT AGGGGGCAGT CTTGCTTTAT TCATCCTCCA	900
	TCTCAAAATG AACTTGGAAT TAAATATTGT AAGATATGTA TAATGCTGGC CATTTTAAAG	960
50	GGGTTTTCTC AAAAGTTAAA CTTTTGTTAT GACTGTGTTT TTGCACATAA TCCATATTTG	1020
	CTGTTCAAGT TAATCTAGAA ATTTATTCAA TTCTGTATGA ACACCTGGAA GCAAAATCAT	1080
5 <i>6</i>	AGTGCAAAAA TACATTTAAG GTGTGGTCAA AAATAAGTCT TTAATTGGTA AATAATAAGC	1140
55	ATTAATTITT TATAGCCTGT ATTCACAATT CTGCGGTACC TTATTGTACC TAAGGGATTC	1200
	TAAAGGTGTT GTCACTGTAT AAAACAGAAA GCACTAGGAT ACAAATGAAG CTTAATTACT	1260
50	AAAATGTAAT TCTTGACACT CTTTCTATAA TTAGCGTTCT TCACCCCCAC CCCCACCCCC	

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	ACCCCCCTTA TTTTCCTTTT GTCTCCTGGT GATTAGGCCA AAGTCTGGGA GTAAGGAGAG	1380
_	GATTAGGTAC TTAGGAGCAA AGAAAGAAGT AGCTTGGAAC TTTTGAGATG ATCCCTAACA	1440
5	TACTGTACTA CTTGCTTTTA CAATGTGTTA GCAGAAACCA GTGGGTTATA ATGTAGAATG	1500
	ATGTGCTTTC TGCCCAAGTG GTAATTCATC TTGGTTTGCT ATGTTAAAAC TGTAAATACA	1560
10	ACAGAACATT AATAAATATC TCTTGTGTAG CACCTTTTAW AAAAAAAAAA AAAAAAAAA	1620
	AAAAAAAA AAAAANCCCG GGGGGGGGCC CCN	1653
15	100	
	(2) INFORMATION FOR SEQ ID NO: 100:	
20	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1145 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 100:	
	TTTTTTTTT TTTTTTTTT TTGACTGAAC TAAGTGGCTT TTTTATTAGA GAAAGCCAGA	60
20	ATTACAAAAG ACTTCCCTTT TCTTGGGGTA TGGCTGTCTC AGCACAATAC TCAACATAAC	120
30	TGCAGAACTG ATGTGGCTCA GGCACCCTGG TTTTAATTCC TTGAGGATCT GGCAATTGGC	180
	TTACGCAAAA GGTCACCATT TGAGGTCCTG CCTTACTAAT TATGTGCTGC CCAACAACTA	240
35	AATTTGTAAT TTGTTTTTCT CTAGTTTGAG CAGGGTCTGA ATTTTTTCAT TTATTTCCTT	300
	TTTTGCCAGC AGACAGACTT GAGTCTGTAA AGACAAGCAA ATACACTGAC AGAAGTTTAC	360
40	CATAGTTTCT AAAATGTAAA AAAGAAAACC CCCAAAAGAC TCAAGAAAAT TAGACCACAA	420
40	ATTITIGCATT GITCATTGTA GCACTATIGG TAATAAAATA ACAAATGTIT GIGCATTITIT	480
	ATGTGAAGAT CCTTCTCGTA TTTCATTTGG AAAGATGAGC AAGAGGTCTG CTTCCTTCAT	540
45	TTTACTTCCC CTTCTGTTTT TGAAAGGCAG TTTCGCCAAG CTTAATGCAA GAATATCTGA	600
	CTGTTTAGAA GAAAGATATT GCCACAATCT CTGGATGGTT TTCCAGGGTT GTGTTATTAC	660
50	TGAGCTTCAT CTTTCCAGAA TGAGCAAAAC ACTGTCCAGT CTTTGTTACG ATTTTGTAAT	720
50	AAATGTGTAC ATTTTTTTA AATTTTTGGA CATCACATGA ATAAAGGTAT GTATGTACGA	780
	ATGTGTATAT ATTATATATA TGACATCTAT TTTGGAAAAT GTTTGCCCTG CTGTACCTCA	840
55	TTTTTAGGAG GTGTGCATGG ATGCAATATA TGAAAATGGG ACATTCTGGA ACTGCTGGTC	900

AGGGGACTTT GTCGCCCTGT GCACTAAAAG GGCCAGATTT TCAGCAGCCA AGGACATCCA

TACCCAAGTG AATGTGATGG GACTTAAAAG AAGTGAACTG AGACAATTCA CTCTGGCTGT

	TTGAACAGCA GCGTTTCATA GGAAGAGAAA AAAAGATCAA TCTTGTATTT TCTGACCACA	1080
	TAAAGGCTTC TTCTCTTTGT AATAAAGTAG AAAAGCTCTC CTCAAAAAAA AAAAAAAAAA	1140
5	AAAAA	1145
••		
10	(2) INFORMATION FOR SEQ ID NO: 101:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 734 base pairs	
15	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 101:	
20	TACCCGGCGG ATTCCAGGAA GGTAAATTTA GTCCTATAAT TTTCAGCTTA ATTATAAACA	60
	AAGGAACAAA TAAGTGGAAG GGCAGCTATT ACCATTCGCT TAGTCAAAAC ATTCGGTTAC	120
25	TGCCCTTTAA TACACTCCTA TCATCAGCAC TTCCACCATG TATTACAAGT CTTGACCCAT	180
	CCCTGTCGTA ACTCCAGTAA AAGTTACTGT TACTAGAAAA TTTTTATCAA TTAACTGACA	240
	AATAGTTTCT TTTTAAAGTA GTTTCTTCCA TCTTTATTCT GACTAGCTTC CAAAATGTGT	300
30	TCCCTTTTTG AATCGAGGTT TTTTTGTTTT GTTTTGTTTT	360
	TGTGCTTCTA TTGCTTTTTT GTGTTTGTT AAGCATGTCC CTTGGCCCAA ATGGAAGAGG	420
35	AAATGTTTAA TTAATGCTTT TTAGTTTAAA TAAATTGAAT CATTTATAAT AATCAGTGTT	480
	AACAATTTAG TGACCCTTGG TAGGTTAAAG GTTGCATTAT TTATACTTGA GATTTTTTTC	540
	CCCTAACTAT TCTGTTTTTT GTACTTTAAA ACTATGGGGG AAATATCACT GGTCTGTCAA	600
40	GAAACAGCAG TAATTATTAC TGAGTTAAAT TGAAAAGTCC AGTGGACCAG GCATTTCTTA	660
	TATAAATAAA ATTGGTGGTA CTAATGTGAA AAAAAAAAAA	720
45	CCGGTACCCT ATTA	734
T J		
50	(2) INFORMATION FOR SEQ ID NO: 102:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 713 base pairs	
	(B) TYPE: nucleic acid	
55	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 102:	
50	CCGCGGGAAC GCTGTCCTGG CTGCCGNCAC CCGAACAGCC TGTCCTGGTG CCCCGGCTCC	60
JU		

	CTGCCCCGCG	CCCAGTCATG	ACCCTGCGCC	CCTCACTCCT	CCCGCTCCAT	CTGCTGCTGC	120
	TGCTGCTGCT	CAGTGCGGCG	GTGTGCCGGG	CTGAGGCTGG	GCTCGAAACC	GAAAGTCCCG	180
5	TCCGGACCCT	CCAAGTGGAG	ACCCTGGTGG	AGCCCCCAGA	ACCATGTGCC	GAGCCCGCTG	240
	CTTTTGGAGA	CACGCTTCAC	ATACACTACA	CGGGAAGCTT	GGTAGATGGA	CGTATTATTG	300
10	ACACCTCCCT	GACCAGAGAC	CCTCTGGTTA	TAGAACTTGG	CCAAAAGCAG	GTGATTCCAG	360
10	GTCTGGAGCA	GAGTCTTCTC	GACATGTGTG	TGGGAGAGAA	GCGAAGGGCA	ATCATTCCTT	420
	CTCACTTGGC	CTATGGAAAA	CGGGGATTTC	CACCATCTGT	CCCAGCGGAT	GCAGTGGTGC	480
15	AGTATGACGT	GGAGCTGATT	GCACTAATCC	GAGCCAACTA	CTGGCTAAAG	CTGGTGAAGG	540
	GCATTTTGCC	TCTGGTAGGG	ATGGCCATGG	TGCCACCCTC	CTGGGCCTCA	TTGGGTATCA	60
20	CCTATACAGA	AAGGCCAATA	GACCCAAAGT	CTCCAAAAAG	AAGCTCAAGG	AAGAGAAACG	66
20	AAACAAGAGC	AAAAAGAAAT	TAATAAATAA	AAATTTTAAA	AAACTTAAAA	AAA	71

30

(2) INFORMATION FOR SEQ ID NO: 103:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1080 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 103:

35 CCGATGTGGA CATCATCCTG TCTATCCCCA TGTTCCTGCG CCTGTACCTG ATCGCCCGAG 60 TCATGCTGCT GCACAGAAGC TCTTCACCGA TGCCTCGTCC CGCAGCATCG GGGCCCTCAA 120 CAAGATCAAC TTCAACACCC GCTTTGTCAT GAAGACGCTC ATGACCATCT GCCCTGGCAC 180 40 TGTGCTGCTC GTGTTCAGCA TCTCTCTGTG GATCATTGCT GCCTGGACCG TCCGTGTCTG 240 TGAAAGTCCT GAATCACCAG CCCAGCCTTC TGGCTCATCA CTTCCTGCTT GGTACCATGA 300 45 360 CCAGCAGGAC GTAACTAGTA ACTTTCTGGG TGCCATGTGG CTCATCTCCA TCACATTCCT TTCCATTGGT TATGGGGACA TGGTGCCCCA CACATACTGT GGGAAAGGTG TCTGTCTCCT 420 CACTGGCATC ATGGGTGCAG GCTGCACTGC CCTTGTGGTG GCCGTGGTGG CCCGAAAGCT 480 50 GGAACTCACC AAAGCGGAGA AGCACGTTCA TAANTTCATG ATGGACACTC AGCTCACCAA 540 GCGGATCAAG AATGYTGCAG CCAATGTCCT TSGGGAAACA TGGTTAATCT ATAAACACAC 600 55 AAAGYTGYTA AAGAAGATTG ACCATGCCAA AGTGAGGAAC ACCAGAGGAA GTTCYTCCAA 660 GTATCCACCA GTTGAGGAGC GTCAAGATGG AACAGAGGAA GCTGAGTGAC CAAGCCAACA 720 780 NICTGGTGGA CCTTTCCAAG ATGCAGAATG TCMTGTATGA CTTAATCACA GAACTCAATG 60

	ACCGGAGCGA AGACCTGGAG AAGCAGATTG GCAGCCTGGA GTCGAAGCTG GAGCATCTCA	840
5	CCGCCAGCTT CAACTCCCTG CCGCTGCTCA TCGCCGACAC CCTGCGCCAG CAGCAGCAGC	900
	AGCTCCTGTC TGCCATCATC GAGGCCCGGG GTGTCAGCGT GGCAGTGGGC ACCACCCACA	960
	CCCCAATCTC CGATAGCCCC ATTGGGGTCA GCTCCACCTC CTTCCCGACC CCGTACACAA	1020
10	GTTCAAGCAG TTGCTAAATA AATCTCCCCA CTCCAGAAGC ATTAAAAAAA AAAAAAAAA	1080
15	(2) INFORMATION FOR SEQ ID NO: 104:	
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 489 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 104:	
25	GGCACGAGAG GCTTTGAAGC ATTTTTGTCT GTGCTCCCTG ATCTTCAGGT CACCACCATG	60
	AAGTTCTTAG CAGTCCTGGT ACTCTTGGGA GTTTCCATCT TTCTGGTCTC TGCCCAGAAT	120
30	CCGACAACAG CTGCTCCAGC TGACACGTAT CCAGCTACTG GTCCTGCTGA TGATGAAGCC	180
	CCTGATGCTG AAACCACTGC TGCTGCAACC ACTGCGACCA CTGCTGCTCC TACCACTGCA	240
	ACCACCGCTG CTTCTACCAC TGCTCGTAAA GACATTCCAG TTTTACCCAA ATGGGTTGGG	300
35	GATCTCCCGA ATGGTAGAGT GTGTCCCTGA GATGGAATCA GCTTGAGTCT TCTGCAATTG	360
	GTCACAACTA TTCATGCTTC CTGTGATTTC ATCCAACTAC TTACCTTGCC TACGATATCC	420
40	CCTTTATCTC TAATCAGTTT ATTTTCTTTC AAATAAAAA TAACTATGAG CAACAAAAA	480
	AAAAAAA	489
45	(2) INFORMATION FOR SEQ ID NO: 105:	
50	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 640 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
55	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 105:	
<i>J J</i>	GCGGTCGCCG CTGTTGTTGT GGTCCCCATG GAGCTGCCGT AGCGGACCCA GCACAGCCAG	60
	GAGCGTCCGG GATGAGCTCA GCCGCGGCCG ACCACTGGGC GTGGTTGCTG GTGCTCAGCT	120
60	TCGTGTTTGG ATGCAATGTT CTTAGGATCC TCCTCCCGTC CTTCTCATCC TTCATGTCCA	180

255

	GGGTGCTGCA	GAAGGACGCG	GAGCAGGAGT	CACAGATGAG	AGCGGAGATC	CAGGACATGA	240
_	AGCAGGAGCT	CTCCACAGTC	AACATGATGG	ACGAGTTTGC	CAGATATGCC	AGGCTGGAAA	300
5	GAAAGATCAA	CAAGATGACG	GATAAGCTCA	AAACCCATGT	GAAAGCTCGG	ACAGCTCAAT	360
	TAGCCAAGAT	AAAATGGGTG	ATAAGTGTCG	CTTTCTACGT	ATTGCAGGCT	GCCCTGATGA	420
10	TCTCACTCAT	TTGGAAGTAT	TATTCTGTCC	CTGTGGCTGT	CGTGCCGAGT	AAATGGATAA	480
	CCCTYTAGAC	CGCCTGGTAG	CCTTTCCYAY	TAGAGTAGCA	GGTGGTGTTG	GAATTACTGT	540
. ~	TGGATTTART	CTGTACAAAT	TGTCCTATTG	TGCTTCACCG	TYCASTGAAC	AGGAGGTGGT	600
15	ACAGCCGGAG	TTAAAAACGG	TTTCCNTTCC	AGTTTAAAAT			640

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25

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(2) INFORMATION FOR SEQ ID NO: 106:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1529 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 106:

GGGCACNAGA TGGAGCTGCC GTAGCGGACC CAGCACAGCC AGGAGCGTCC GGGATGAGCT 60 CAGCCGCGC CGACCACTGG GCGTGGTTGC TGGTGCTCAG CTTCGTGTTT GGATGCAATG 120 TTCTTAGGAT CCTCCTCCCG TCCTTCTCAT CCTTCATGTC CAGGGTGCTG CAGAAGGACG 180 35 CGGACAGGAG TCACAGATGA GAGCGGAGAT CCAGGACATG AAGCAGGAGC TCTCCACAGT 240 CAACATGATG GACGAGTTTG CCAGATATGC CAGGCTGGAA AGAAAGATCA ACAAGATGAC 40 GGATAAGCTC AAAACCCATG TGAAAGCTCG GACAGCTCAA TTAGCCAAGA TAAAATGGGT 360 GATAAGTGTC GCTTTCTACG TATTGCAGGC TGCCCTGATG ATCTCACTCA TTTGGAAGTA 420 TTATTCTGTC CCTGTGGCTG TCGTGCCGAG TAAATGGATA ACCCCTCTAG ACCGCCTGGT 480 45 AGCCTTTCCT ACTAGAGTAG CAGGTGGTGT TGGAATTACC TGTTGGATTT TAGTCTGTAA 540 CAAAGTTGTC GCTATTGTGC TTCATCCGTT CAGCTGAACA GGAGGATGGA TACAGCCGCG 600 50 AGTAAAAAA CGGATTTCCT CTTCCTAGCT TAAAATCTGA TTTACACTGT TTTGTTTTTT 660 AAGAAACAAA AGTGCATAGT TTAGATTTTT TTTTTGTTGA ATATGTTTGT TCTTGGACTT 720 TATGAGATAG TCTTATAAGA ATCACGATTT TCTACACCTG TCATTGAGCC AAGAAAGTCC 780 55 AGTTTATGAC ACGTATGTAC TAGTGAACAC CGTCCTCGAT CTGTACGAAA TGTGAAATGT 840 900 TTAGGGACAT CTCCATGCTG TCACTTGTGA TTTGCCCTCT TATGTATTTT GGTCATATTG 60

	CCAACTGGAA	AGTCAAAATT	TTCTAACAAC	TTTAAGTAAG	TTCTTTGAAG	ACTTAGTGCT	960
	GTTTTTAATC	CAGTTTAGAA	AGTAACTTAA	TTTTAATACC	RCTACTAAAA	ATTCGAAAAT	1020
5	TTCTTCTTTA	ATCACATTCA	ATATGGTTAA	AAGAACAACA	CTAATTGACA	TTGCGTGGGC	1080
	TTTTTCTCCC	TTTGTTTAAA	ATGTCATTTG	TTGAGCAAGA	GTTGTATAGT	ATTATCTACT	1140
10	TACTTGAGGC	TGTTAATTTT	TCATTACAGT	GTTTTGTAAA	TGTATCCACG	AGACCATGAT	1200
	GCATTGTTTT	GTGCTCAACT	TGTGTTTTGT	ATTTAAAGCA	TTTTGAATGA	AGTGTATTTT	1260
	ATAAGCATTT	AATATTTATG	CTCTTTAGAA	TGGAACACAG	AAAACAAACC	TTATAAGTCC	1320
15	TGATTAATCT	GAACCAATAA	CCTGTGTGGC	CTACAAAGTA	TAATTCTATT	AAATGTTCCT	1380
	TAAAACACTT	TTTTCTAATT	AAAATCTTTG	CAAATGCTTG	TGTAACTTCC	TGCCTTACAG	1440
20	CTACTTGTTT	GCTGTGAGCC	ACCCGCAACT	GACAAGTGGC	TGTTAACTGA	GTCACCATAT	1500
	CCCAGTAAAG	CTGAATTTTC	TCACTAAAA				1529
25	(2) INFORMA	ATION FOR SE	CO ID NO. 10	17 •			
30	(1)		HARACTERISTI GTH: 2435 ba E: nucleic a	ase pairs			

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 107: 35

ATGAAGGGTC GTTGGTGGGA AAGATGGCGG CGACTCTGGG ACCCCTTGGT CGTGGCAGCA 60 GTGGCGRCGA TGTTTGTCGG CTCGGGATGG GTCCAGGATG TTACTCCTTC TTCTTTTGTT 120 40 GGGGTCTGGG CAGGGGCCAC AGCAAGTCGG GGCGGGTCAA ACGTTCGAGT ACTTGAAACG 180 GGAGCACTCG CTGTCGAAGC CCTACCAGGG TGTGGGCACA GGCAGTTCCT CACTGTGGAA 240 TCTGATGGGC AATGCCATGG TGATGACCCA GTATATCCGC CTTACCCCAG ATATGCAAAG 300 45 TAAACAGGGT GCCTTGTGGA ACCGGGTGCC ATGTTTCCTG AGAGACTGGG AGTTGCAGGT 360 GCACTTCAAA ATCCATGGAC AAGGAAAGAA GAATCTGCAT GGGGATGGCT TGGCAATCTG 420 50 GTACACAAAG GRWTCGGATG CAGCCAGGGC CTGTNTTTGG GAAACATGGA CAAATTTGTG 480 GGGCTGGGAG TATTTGTAGA CACCTACCCC AATGAGGAGA AGCAGCAAGA GCGGGTATTC 540 CCCTRCMTCT CAGCCATGGT GAACAACGGC TCCCTCAGCT ATGATCATGA GCGGGATGGG 600 55 CGGCCTACAG AGCTGGGAGG CTGCASAGCC ATTGTCCGCA ATCTTCATTA CGACACCTTC 660 CTGGTGATTC GCTACGTCAA GAGGCATTTR ACGATAATGA TGGATATTGA TGGCAAGCAT 720 60 GAGTGGAGGG ACTGCATTGA AGTGCCCGGA GTCCGCCTGC CCCGCGGCTA CTACTTCGGC 780

	ACCTCCTCCA TCACTGGGGA TCTCTCAGAT AATCATGATG TCATTTCCTT GAAGTTGTTT	840
_	GAACTGACAG TGGAGAGAAC CCCAGAAGAG GAAAAGCTCC ATCGAGATGT GTTCTTGCCC	900
5	TCAGTGGACA ATATGAAGCT GCCTGAGATG ACAGCTCCAC TGCCGCCCCT GAGTGGCCTG	960
	GCCCTCTTCC TCATCGTCTT TTTCTCCCTG GGTGTTTTCT GTATTTGCCA TAGTCATTGG	1020
10	TATCATACTC TACAACAAAT GGCAGGAACA GAGCCGAAAG CGCTTCTACT GAGCCCTCCT	1080
	GCTGCCACCA CTTTTGTGAC TGTCACCCAT GAGGTATGGA AGGAGCAGGC ACTGGCCTGA	1140
15	GCATGCAGCC TGGAGAGTGT TCTTGTCTCT AGCAGCTGGT TGGGGACTAT ATTCTGTCAC	1200
13	TGGAGTTTTG AATGCAGGGA CCCCGCATTC CCATGGTTGT GCATGGGGAC ATCTAACTCT	1260
	GGTCTGGGAA GCCACCCACC CCAGGGCAAT GCTGCTGTGA TGTGCCTTTC CCTGCAGTCC	1320
20	TTCCATGTGG GAGCAGAGGT GTGAAGAGAA TTTACGTGGT TGTGATGCCA AAATCACAGA	1380
	ACAGAATTTC ATAGCCCAGG CTGCCGTGTT GTTTGACTCA GAAGGCCCCTT CTACTTCAGT	1440
25	TTTGAATCCA CAAAGAATTA AAAACTGGTA ACACCACAGG CTTTCTGACC ATCCATTCGT	1500
23	TGGGTTTTGC ATTTGACCCA ACCCTCTGCC TACCTGAGGA GCTTTCTTTG GAAACCAGGA	1560
	TGGAAACTTC TTCCCTGCCT TACCTTCCTT TCACTCCATT CATTGTCCTC TCTGTGTGCA	1620
30	ACCTGAGCTG GGAAAGGCAT TTGGATGCCT CTCTGTTGGG GCCTGGGGCT GCAGAACACA	1680
	CCTGCGTTTC ACTGGCCTTC ATTAGGTGGC CCTAGGGAGA TGGCTTTCTG CTTTGGATCA	1740
35	CTGTTCCCTA GCATGGGTCT TGGGTCTATT GGCATGTCCA TGGCCTTCCC AATCAAGTCT	1800
55	CTTCAGGCCC TCAGTGAAGT TTGGCTAAAG GTTGGTGTAA AAATCAAGAG AAGCCTGGAA	1860
	GACATCATGG ATGCCATGGA TTAGCTGTGC AACTGACCAG CTCCAGGTTT GATCAAACCA	1920
40	AAAGCAACAT TTGTCATGTG GTCTGACCAT GTGGAGATGT TTCTGGACTT GCTAGAGCCT	1980
	GCTTAGCTGC ATGTTTTGTA GTTACGATTT TTGGAATCCC ACTTTGAGTG CTGAAAGTGT	2040
45	AAGGAAGCTT TCTTCTTACA CCTTGGGCTT GGATATTGCC CAGAGAAGAA ATTTGGCTTT	2100
73	TTTTTTNCTT AATGGACAAG AGACAGTTGC TGTTCTCATG TTCCAAGTCT GAGAGCAACA	2160
	GACCCTCATC ATCTGTGCCT GGAAGAGTTC ACTGTCATTG AGCAGCACAG CCTGAGTGCT	2220
50	GGCCTCTGTC AACCCTTATT CCACTGCCTT ATTTGACAAG GGGTTACATG CTGCTCACCT	2280
	TACTGCCCTG GGATTAAATC AGTTACAGGC CAGAGTCTCC TTGGAGGGCC TGGAACTCTG	2340
55	AGTCCTCCTA TGAACCTCTG TAGCCTAAAT GAAATTCTTA AAATCACCGA TGGAACCAAA	2400
JJ	MAAAA AAAAAAAAA AAAAAAAAA AAAAAAAAA	2435

	(2) INFORMATION FOR SEQ ID NO: 108:	
5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 805 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 108:	
	ATGAAACTTA AGAATTGAAT TGGAAAGACT TCTCAAAGAG AATTGTATGT AACGATGTTG	60
	TATTGATTTT TAAGAAAGTA ATTTAATTTG TAAAACTTCT GCTCGTTTAC ACTGCACATT	120
15	GAATACAGGT AACTAATTGG AAGGAGAGGG GAGGTCACTC TTTTGATGGT GGCCCTGAAC	180
	CTCATTCTGG TTCCCTGCTG CGCTGCTTGG TGTGACCCAC GGAGGATCCA CTCCCAGGAT	240
20	GACGTGCTCC GTAGCTCTGC TGCTGATACT GGGTCTGCGA TGCAGCGGCG TGAGGCCTGG	300
	GCTGGTTGGA GAAGGTCACA ACCCTTCTCT GTTGGTCTGC CTTCTGCTGA AAGACTCGAG	360
	AACCAACCAG GGAAGCTGTC CTGGAGGTCC CTGGTCGGAG AGGGACATAG AATCTGTGAC	420
25	CTCTGACAAC TGTGAAGCCA CCCTGGGCTA CAGAAACCAC AGTCTTCCCA GCAATTATTA	480
	CAATICTTGA ATTCCTTGGG GATTTTTTAC TGCCCTTTCA AAGCACTTAA GTGTTAGATC	540
30	TAACGTGTTC CAGTGTCTGT CTGAGGTGAC TTAAAAAATC AGAACAAAAC TTCTATTATC	600
	CAGAGTCATG GGAGAGTACA CCCTTTCCAG GAATAATGTT TTGGGAAACA CTGAAATGAA	660
	ATCTTCCCAG TATTATAAAT TGTGTATTTA AAAAAAAGAA ACTTTTCTGA ATGCCTACTG	720
35	GCGGTGTATA CCAGGCAGTG TGCCAGTTTA AAAAGATGAA AAAGAATAAA AACTTTTGAG	780
	GAACAAAAA AAAAAAAAA AAATT	805
40		
	(2) INFORMATION FOR SEQ ID NO: 109:	
45	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1166 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 109:	
	GGCACGAGAG GCGCCAGTCG CAGGTGTGCT GCTGAGGCGT GAGAATGGCG TCCCGCGGCC	60
55	GGCGTCCGGA GCATGGCGGA CCCCCAGAGC TGTTTTATGA CGAGACAGAA GCCCGGAAAT	120
	ACGTTCGCAA CTCACGGATG ATTGATATCC AGACCAGGAT GGCTGGGCGA GCATTGGAGC	180
	TTCTTTATCT GCCAGAGAAT AAGCCCTGTT ACCTGCTGGA TATTGGCTGT GGCACTGGGC	240

TGAGTGGAAG TTATCTGTCA GATGAAGGGC ACTATTGGGT GGGCCTGGAT ATCAGCCCTG

	CCATGCTGGA	TGAGGCTGTG	GACCGAGAGA	TAGAGGGAGA	CCTGCTGCTG	GGGGATATGG	360
ے	GCCAGGGCAT	CCCATTCAAG	CCAGGCACAT	TTGATGGTTG	CATCAGCATT	TCTGCTGTGC	420
5	AGTGGCTCTG	TAATGCTAAC	AAGAAGTCTG	AAAACCCTGC	CAAGCGCCTG	TACTGCTTTT	480
	TIGCTICTCT	TTTTTCTGTT	CTCGTCCGGG	GATCCCGAGC	TGTCCTGCAG	CTGTACCCTG	540
10	AGAACTCAGA	GCAGTTGGAG	CTGATCACAA	CCCAGGCCAC	AAAGGCAGGC	TTCTCCGGTG	600
	GCATGGTGGT	AGACTACCCT	AACAGTGCCA	AAGCAAAGAA	ATTCTACCTC	TGCTTGTTTT	660
	CTGGGCCTTC	GACCTTTATA	CCAGAGGGGC	TGAGTGAAAA	TCAGGATGAA	GTTGAACCCA	720
15	GGGAGTCTGT	GTTCACCAAT	GAGAGGTTCC	CATTAAGGAT	GTCGAGGCGG	GGAATGGTGA	780
	GGAAGAGTCG	GGCATGGGTG	CTGGAGAAGA	AGGAGCGGCA	CAGGCGCCAG	GGCAGGGAAG	840
20	TCAGACCTGA	CACCCAGTAC	ACCGGCCGCA	AGCGCAAGCC	CCGCTTCTAA	GTCACCACGC	900
	GGTTCTGGAA	AGGCACTTGC	CTCTGCACTT	TTCTATATTG	TTCAGCTGAC	AAAGTAGTAT	960
~ ~	TTTAGAAAAG	TTCTAAAGTT	ATAAAAATGT	TTTCTGCAGT	DAAAAAAA	TTCTCTGGGC	1020
25	CGGGCGTGGT	GGCTCACANC	TGTAATCCCA	GCACCTTGGG	; AGGCTGAGGT	GGGAGGATCA	1080
	TTTGAGGCCA	A GGAGTTTGAG	ACCTGCCTGC	GCAACATAA1	GAAACTTCCT	TTCCAGGGAG	1140
30	AAAAAAAA	AAAAAAAA A	ACTCGA				1166

35 (2) INFORMATION FOR SEQ ID NO: 110:

WO 98/42738

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 586 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 110:

60 AGAGCGGACG AAGCTGGATA ACAGGGGACC GATGATGTGG CGACCATCAG TTCTGCTGCT 45 TCTGTTGCTA CTGAGGCACG GGGCCCAGGG GAAGCCATCC CCAGACGCAG GCCCTCATGG 120 CCAGGGGAGG GTGCACCAGG CGGCCCCCCT GAGCGACGCT CCCCATGATG ACGCCCACGG 180 50 GAACTTCCAG TACGACCATG AGGCTTTCCT GGGACGGGAA GTGGCCAAGG AATTCGACCA 240 ACTCACCCCA GAGGAAAGCC AGGCCCGTCT GGGGCGGATC GTGGACCGCA TGGACCGCGC 300 GGGGGACGGC GACGGCTGGG TGTCGCTGGC CGAGCTTCGC GCGTGGATCG CGCACACGCA 360 55 420 GCAGCGGCAC ATACGGGACT CGGTGAGCGC GGCCTGGGAC ACGTACGACA CGGACCGCGA CGGGCGTGTG GGTTGGGAGG AGCTGCGCAA CGYCACCTAT GGCCACTASG SGCCCGKTGA 480 60

	AGAATTTCAT GACGTGGAGG ATGCAGAGAC YTACAAAAAG ATGCTGGYTC GGGACGAGCG	540
	GCGTTTCCGG GTGGCCGACC AGGATGGGGA CTCGATGGCC ACTCGA	586
5		
	(2) INFORMATION FOR SEQ ID NO: 111:	
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 1134 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 111:	
	ACCCATTGAG CAGAAGGAGG CCAGGTGGGA AAGCTCCTGG GAAGAGCAGC CAGACTGGAC	60
20	ACTGGGCTGC TTGAGTCCTG AGTCACAATT CAGAATTCCT GGGCTCCCTG GGTGCATTCT	120
	ATCATTCCAG TTGAAAGTTT GCTTCCTTCC AGTCATGTGG CTCTTCATTC TACTCTCCTT	180
25	GGCTCTCATT TCAGATGCCA TGGTCATGGA TGAAAAGGTC AAGAGAAGTT TGTGCTGGAC	240
	ACGGCTTCTG CCATCTGCAA CTACAATGCC CAYTACAAGA ATCACCCCAA ATACTGGTGC	300
	CGAGGYTATT TCCGTGAYTA CTGCAACATC ATCGCCTTCT CCCCTAACAG CACCAATCAT	360
30	GTGGCCCTGA AGGACACAGG GAACCAGCTC ATTGTCACTA TGTCCTGCCT GAACAAANAA	420
	GACACGGGCT GGTACTGGTG TGGCATCCAR CGGGACTTTG CMAGGGATGA CATGGATTTT	480
35	ACAGAGCTGA TTGTAACTGA CGACAAAGGA ACCCTGGCCA ATGACTTTTG GTCTGGGAAA	540
	GACCTATCAG GCAACAAAAC CAGAAGCTGC AAGGCTCCCA AAGTTGTCCG CAAGCTGACC	600
	GCTCCAGGAC GTCCATTCTC ATCATTTGCA TACTGATCAC GGGTTTGGGA ATCATCTCTG	660
40	TAATCAGTCA TTTGACCAAA AGGAGGAGAA GTCAAAGGAA TAGAAGGGTA GGCAACACTT	720
	TGAAGCCCTT CTCGCGTGTC CTGACTCCAA AGGAAATGGC TCCTACTGAA CAGATGTGAC	780
45	TGAAGWITTT TTTAATTTAG TTNCATAAAG TGATGNCTAC AACAGAWTAA TCACCCATGA	840
	CAACTGGCCC CACACCTCAG AGACTGATTC TGATCTCCCA GGAATTCTGA AGGACCCTCT	900
	ATCCTTGACA ACAATCATTT GCAGCCAGGT AGCAACGGCR GTAGTCAGAG GAGCTATGAT	960
50	AGACCACACC CAAGCAAGGC TGCCCTCAAA TAACATCTCA AGATCTTAGT TCTTATGCAT	1020
	TCCATCAGTC AGAAGTGAAG AAGAGGTGGA GAATCTKGAT TGGGGACCAG GAAATCACTT	1080
55	GTATTTTGTT AGCCAATAAA TTCCTAGCCA GTGTTGAATG AAAAAAAAAA	1134

(2) INFORMATION FOR SEQ ID NO: 112:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1333 base pairs(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

5 (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 112:

10	CACTITAAAG CTCTGCTGAG GGAGTTCGGA GCCCAGGCTT TCAGGCGACC TCTGCCCTCC	60
10	CTGCCTCTCC TCACCCTCCC TCTCTTCCTG CAGGGCCTGG GAAGGGCTTT GAGGGAGCCT	120
	GGGAGCCATG TGAAGAGGGG CACGCCTGGG CTGTCCCACA GTTTAGATCC AGTTGGAGGT	180
15	TCTCCCTGGC TCCTGCAGGC CTGCGGGGAT CTCTCCCCAC TTCAGGCCTC CGGCAGCTGC	240
	CTGCCCTCTT GTCTGTGCTT CAGCCCTGCA CAAAAGCAGC TTGGTGACAC CACTCAGCCA	300
20	CCCAGAGTAC GTGTTTACAG GCTTTCCAGA TCACCTTCCT GTGGGGTGAA CGTAATGAGG	360
20	CGGGGCTGGT CCTTGGAATT TCCCCTGGAA AATGGTAACA GACTCCATCC TTGACCCGGG	420
	GATGAGCATG AAGGCATTGT CCCAAAGGCA GAGGCCACCG TGGTAGGAAT TCCACCAAGG	480
25	CCAGAAGGGA AAAAGGAAGA ACCCACCGTG TCTGGCTGTG CGGGCCCTGG GGAGGGTCGT	540
	GAGTGCAGCC CCTCTCTACT TCYGTGCCTT TGTAAAACGT GTAGATAACC GCAGTGGTTG	600
30	GCTGAGCCAA GAACTCTCCT AAATCAGTGG CTTTCTCCCC ACCCCTTGCT GGGGAGTCAT	660
30	TTTTAAAAAA ATCTGTGGGA TATAAAATTG GCCTCCTGCT GCTTCAGCCT ACCTCTCCCT	720
	CTGCTGACTT AATGTCGTGA TTCTGTTTCT TCAGATATTT AAGGCTGTTA GGTTGTGTGA	780
35	GCCTTGAAGT GTGTGTGTG GTCCCAGCGA CTGTCCACTG TCCAGGAGAT GCATGTCTTT	840
	GTATTGGAGA TATTTCTGTA ACTCATTCTC TTGGTGCTCA CGATTGCCAT GGCCATAGGG	900
40	CCACAGTGCC GTATCTGCTG CAGACATGAT TGTTTCTTGT TCTAGAGGTT TTCTTGTTTT	960
40	CGAATCTTGC CTGATGAATC CAGCCAGACC AAGGGGCCTA GATTTGACCT CTGTCCTGGG	1020
	CTCCTGGGCC AGGTGCAGGA ACATCTGAGG CCACTCTGCT GGCCACCTCC AGTGGGTGCT	1080
45	GACCACAGGA TGGGCTTTGT TTACACTCAT TTTCACCCTG ATTCTTGCCC CCACTTTCAT	1140
	AAAAGAAACT TCAAAATGCT GACGCTTTGG AGAGTAAGAA AATCAATCTT GGCTGGGCAC	1200
50	GGTGGCTCCT GCCTGTGATC CTAGCACTTT GGGAGGCTGA AGCTGAAGGA TCACTTGAGC	1260
50	TCAGGAGTTG GAGACCAACC CTGGCAACAT AACAAGACCC TGTCTCTACA AAAAAAAAAA	1320
	AAAAAAAACT CGA	1333

(2) INFORMATION FOR SEQ ID NO: 113:

(i) SEQUENCE CHARACTERISTICS:

55

5	(A) LENGTH: 1015 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 113:	
10	GGCACGAGCG GCACGAGCGG CACGAGGTGA CTTCAAGTGT CGGATCTTTT CAGCCTACAT	60
10	CAAGGAGGTG GAGGAACGGC CGGCACCCAC CCCGTGGGCT CCAAGATGCC CTTTGGGGAA	120
	CTGATGTTCG AATCCAGCAG TAGCTGCGGC TGGGTACATG GCGTCTGTTT CTCAGCCAGC	180
15	GGGAGCCGCG TGGCCTGGGT AAGCCACGAC AGCACCGTCT GCCTGGCTGA TGCCGACAAG	240
	AAGATGGCCG TCGCGACTCT GGCCTCTGAA ACACTACCAC TGCTGGCGCT GACCTTCATC	300
	ACAGACAACA GCCTGGTGGC AGCGGGCCAC GACTGCTTCC CGGTGCTGTT CACCTATGAC	360
20	GCCGCCGCGG GGATGCTGAG CTTCGGCGGG CGGCTGGACG TTCCTAAGCA GAGCTCGCAG	420
	CGTGGCTTGA CGGCCCGCGA GCGCTTCCAG AACCTGGACA AGAAGGCGAG CTCCGAGGGT	480
25	GGCACGGCTG CGGGCGCGGG CCTAGACTCG CTGCACAAGA ACAGCGTCAG CCAGATCTCG	540
	GTGCTCAGCG GCGGCAAGGC CAAGTGCTCG CAGTTCTGCA CCACTGGCAT GGATGGCGGC	600
	ATGAGTATCT GGGATGTGAA GAGCTTGGAG TCAGCCTTGA AGGACCTCAA GATCAAATGA	660
30	CCTGTGAGGA ATATGTTGCC TTCATCCTAG CTGCTGGGGA AGCGGGGAGA GGGGTCAGGG	720
	AGGCTAATGG TTGCTTTGCT GAATGTTTCT GGGGTACCAA TACGAGTTCC CATAGGGGCT	780
35	GCTCCCTCAA AAAGGGAGGG GACAGATGGG GAGCTTTTCT TACCTATTCA AGGAATACGT	840
	GCCTTTTTCT TAAATGCTTT CATTTATTGA AAAAAAAAA AAATGCCCCC AAAGCACTAT	900
	GCTGGTCATG AACTGCTTCA AAATGTGGAG GTAATAAAAT GCAACTGTGT AAAAAAAAAA	960
40	ААААААААА ААААААААА ААААААААА АААААААА	1015
45	(2) INFORMATION FOR SEQ ID NO: 114:	
50	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 1076 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 114:	
55	GGCACGAGGG GAAAGCCATG CTCCCAGGAC TCCTTCCTTG CAGCCTTAAA TCGGTCTGTA	60
	CGGAAAATTC CGCGCCTTAG AAACCCACGC TTGGGTGTAA CTTATTATTG TTCTTCCTGA	120
50	CCTACTTCCT GTTTATCACT TCCGGGTTCA TCATTTTGGC ATTTCGGTGA TCGGGTTGGA	180
50		

	ACTATTGAAG CCCGCTTTCA GGTTCTTTTC CCCATTTTCC CTTTGAAAGG AAGACTTCTG	240
	GCTTCTCCTA AATCTCCGTT CTCTGGGTAA GGGGAGTCCA AGCCTCTGTC ATGAGGAACG	300
5	GAAATGCGAG GGCCTCGGGT GTTACTCTAA AATCCGCCCT CAGCTTGCAC GCCGGAAGCT	360
	GCGATTCCTG CAGCGGAAGA GGCGTGATCT GGCCTTCGAC TCGCTATGTC CACTAACAAT	420
	ATGTCGGACC CACGGAGGCC GAACAAAGTG CTGAGGTACA AGCCCCCGCC GAGCGAATGT	480
10	AACCCGGCCT TGGACGACCC GACGCCGGAC TACATGAACC TGCTGGGCAT GATCTTCAGC	540
	ATGTGCGGCC TCATGCTTAA GCTGAAGTGG TGTGCTTGGG TCGCTGTCTA CTGCTCCTTC	600
15	ATCAGCTTTG CCAACTCTCG GAGCTCGGAG GACACGAAGC AAATGATGAG TAGCTTCATG	660
13	CTGTCCATCT CTGCCGTGGT GATGTCCTAT CTGCAGAATC CTCAGCCCAT GACGCCCCCA	720
	TGGTGATACC AGCCTAGAAG GGTCACATTT TGGACCCTGT CTATCCACTA GGCCTGGGCT	780
20	TTGGCTGCTA AACCTGCTGC CTTCAGCTGC CATCCTGGAC TTCCCTGAAT GAGGCCGTCT	840
	CGGTGCCCC AGCTGGATAG AGGGAACCTG GCCCTTTCCT AGGGAACACC CTAGGCTTAC	900
25	CCCTCCTGCC TCCCTTCCCC TGCCTGCTGC TGGGGGAGAT GCTGTCCATG TTTCTAGGGG	960
43	TATTCATTTG CTTTCTCGTT GAAACCTGTT GTTAATAAAG TTTTTCACTC TGAAAAAAAA	1020
	AAAAAAAANA RAAAACNCGN GGGGGGGCCC GGAACCCAAT TCSCCGGATA GTGAGT	1076
30	AMMAMMAN INMACREON COCCOCCCC COLUMN TO THE TOTAL	
35	(2) INFORMATION FOR SEQ ID NO: 115:	
33	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1487 base pairs	
	(B) TYPE: nucleic acid	
40	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 115:	
4.5	CCGCTGCTGA TAACTATGGC ATCCCCCGGG CCTGCAGGAA TTCGGCACGG AGCTACGGCG	60
45	CCGCCTGGCT CCTGCTGNCA CCTGCAGGCT CGTCGCGGGT GGAGCCCACC CAAGACATCA	120
	GCATCAGCGA CCAGCTGGGG GGCCAGGACG TGCCCGTGTT CCGGAACCTG TCCCTGCTGG	180
50	TGGTGGGTGT CGGCGCCGTG TTCTCACTGC TATTCCACCT GGGCACCCGG GAGAGGCGCC	240
	GGCCGCATGC GGASGAGCCA GGCGAGCACA CCCCCCTGTT GGCCCCTGCC ACGGCCCAGC	300
ے ہے	CCCTGCTGCT CTGGAAGCAC TGGCTCCGGG AGCSGGCTTT CTACCAGGTG GGCATACTGT	360
55	ACATGACCAC CAGGCTCATC GTGAACCTGT CCCAGACCTA CATGGCCATG TACCTCACCT	420
	ACTICIONAL COTOCOCA A ACTITICATICA COACCATITUD COTOCITGATG TACCITCAGCG	480

GCTTCTTGTC CTCCTTCCTC ATGAAGCCCA TCAACAAGTG CATTGGGAGG AACATGACCT

	ACTTCTCAGG	CCTCCTGGTG	ATCCTGGCCT	TTGCCGCCTG	GGTGGCGCTG	GCGGAGGGAC	600
5	TGGGTGTGGC	CGTGTACGCA	GCGGCTGTGC	TGCTGGGTGC	TGGCTGTGCC	ACCATCCTCG	660
	TCACCTCGCT	GGCCATGACG	GCCGACCTCA	TCGGTCCCCA	CACGAACAGC	GGAGCKTTCG	720
	TGTACGGCTC	CATGAGCTTC	TTGGATAAGG	TGGCCAATGG	GCTGGCAGTC	ATGGCCATCC	780
10	AGAGCCTGCA	CCCTTGCCCC	TCAGAGCTCT	GCTGCAGGGC	CTGCGTGAGC	TTTTACCACT	840
	GGGCGATGGT	GGCTGTGACG	GGCGGCGTGG	GCGTGGCCGC	TGCCCTGTGT	CTCTGTAGCC	900
15	TCCTGCTGTG	GCCGACCCGC	CTGCGACGCT	GATGAGACCT	GCACGCANTG	GCTCACAGCA	960
	GCACGATTTG	TGACAGCCCG	AGGCGGAGAA	CACCGAACAC	CCAGTGAAGG	TGAGGGGATC	1020
	AGCACGGCGC	GGCCACCCAC	GCACCCACGC	GCTGGAATGA	GACTCAGCCA	CAAGGAGGTG	1080
20	CGAAGCTCTG	ACCCAGGCCA	CAGTGCGGAT	GCACCTTGAG	GATGTCACGC	TCAGTGAGAG	1140
	ACACCAGACA	CAGAAGGGTA	CGCTGTGATC	CCACTTCTAT	GAAATGTCCA	GGACAGACCA	1200
25	ATCCACAGAA	TCAGGGAGAG	GATTCGTGGG	TGCCGGGACT	GGGGAGGGGG	ACCTGGGGGT	1260
	GACTAGGTGA	CATAATGGGG	ACAGGGCTGC	CTTCTGGGTG	ATGAGAATGT	TCTGGAATCA	1320
	GATGGGATGG (CTGCACGGCG	TGGTGAAGGT	ACTGAACGCC	ACCTCACTGT	AAGACGGTAG	1380
30	ATTTTGTATT	TTACCACAAT	AAACAAAACA	AAACAAAACC	AAAAAAAAA	AAAAAAAA	1440
	AAAAAAAAGG 2	AATTCGATAT	CAAGCTTATC	GATACCGTCG	ACCTCGA		1487
35							
	(2) INFORMA	TION FOR SE	Q ID NO: 11	6:			
40	(i)	(A) LENC (B) TYPE (C) STRA	WARACTERISTI FTH: 1350 ba E: nucleic a ANDEDNESS: C DLOGY: linea	ase pairs acid double			
45	(xi)	SEQUENCE D	ESCRIPTION:	SEQ ID NO:	116:		
	GGCACGAGTG (CGCANGCGTG	GGGCTCTCTC	CTTGTCAGTC	GCCCCCCT	GCGGGCTGGT	60
50	GGCTCTGTGG (CAGCGGCGGC	GGCAGGACTC	CGGCACTATG	AGCGGCTTCA	GCACCGAGGA	120
30	GCGCGCGCG (CCTTCTCCCT	GGAGTACCGA	GTCTTCCTCA	Aaaatgagaa	AGGACAATAT	180
	ATATCTCCAT 1	TCATGATAT '	TCCAATTTAT	GCAGATAAGG	ATGTGTTTCA	CATGGTAGTT	240
55	GAAGTACCAC	GCTGGTCTAA '	TGCAAAAATG	GAGATTGCTA	CAAAGGACCC	TTTAAACCCT	300
	ATTAAACAAG A	ATGTGAAAAA .	AGGAAAACTT	CGCTATGTTG	CGAATTTGTT	CCCGTATAAA	360
50	GGATATATCT C	GAACTATGG '	IGCCATCCCT	CAGACTTGGG	AAGACCCAGG	GCACAATGAT	420

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	AAACATACTG GCTGTTGTGG TGACAATGAC CCAATTGATG TGTGTGAAAT TGGAAGCAAG	480
	GTATGTGCAA GAGGTGAAAT AATTGGCGTG AAAGTTCTAG GCATATTGGC TATGATTGAC	540
5	GAAGGGGAAA CCGACTGGAA AGTCATTGCC ATTAATGTGG ATGATCCTGA TGCAGCCAAT	600
	TATAATGATA TCAATGATGT CAAACGGCTG AAACCTGGCT ACTTAGAAGC TACTGTGGAC	660
10	TGGTTTAGAA GGTATAAGGT TCCTGATGGA AAACCAGAAA ATGAGTTTGC GTTTAATGCA	720
10	GAATTTAAAG ATAAGGACTT TGCCATTGAT ATTATTAAAA GCACTCATGA CCATTGGAAA	780
	GCATTAGTGA CTAAGAAAAC GAATGGAAAA GGAATCAGTT GCATGAATAC AACTTTGTCT	840
15	GAGAGCCCCT TCAAGTGTGA TCCTGATGCT GCCAGAGCCA TTGTGGATGC TTTACCACCA	900
	CCCTGTGAAT CTGCCTGCAC AGTACCAACA GACGTGGATA AGTGGTTCCA TCACCAGAAA	960
20	AACTAATGAG ATTTCTCTGG AATACAAGCT GATATTGCTA CATCGTGTTC ATCTGGATGT	1020
20	ATTAGAAGTA AAAGTAGTAG CTTTTCAAAG CTTTAAATTT GTAGAACTCA TCTAACTAAA	1080
	GTAAATTCTG CTGTGACTAA TCCAATATAC TCAGAATGTT ATCCATCTAA AGCATTTTTC	1140
25	ATATCTCAAC TAAGATAACT TTTAGCACAT GCTTAAATAT CAAAGCAGTT GTCATTTGGA	1200
	AGTCACTTGT GAATAGATGT GCAAGGGGAG CACATATTGG ATGTATATGT TACCATATGT	1260
30	TAGGAAATAA AATTATTTTG CTGAAAAAAA AAAAAAAAA ACCTSGGGG GGGSCCCGGT	1320
30	CCCCATTTGG CCCTTTGGGG GGNGGTTTTA	1350
35	(2) INFORMATION FOR SEQ ID NO: 117:	
	(i) SEQUENCE CHARACTERISTICS:	
40	(A) LENGTH: 2527 base pairs	
40	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
	(D) TOPOLOGY: linear	
45	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 117:	
	CTCTTGCTAC CTTCCCGGCG CAGAGAACCC CGGCTGCTCA GCGCGCTCCG GGGTCATGGA	60
	GATCCCCGGG AGCCTGTGCA AGAAAGTCAA GCTGAGCAAT AACGCGCAGA ACTGGGGAAT	120
50	GCAGAGAGCA ACCAATGTCA CCTACCAAGC CCATCATGTC AGCAGGAACA AGAGAGGTCA	180

GGTGGTGGGG ACCAGAGGTG GCTTTCGTGG TTGCACAGTT TGGCTAACAG GCTTGTCTGG

AGCGGGAAAG ACTACTGTGA GCATGGCCTT GGAGGAGTAC CTGGTTTGTC ATGGTATTCC

ATGCTACACT CTGGATGGTG ACAATATTCG TCAAGGTCTC AATAAAAATC TTGGCTTTAG

TCCTGAAGAC AGAGAAGAGA ATGTTCGACG CATCGCAGAA GTTGCTAAAC TGTTTGCAGA

TGCTGGCTTA GTGTGCATCA CAAGTTTCAT ATCACCTTAC ACTCAGGATC GCAACAATGC

240

300

360

420

480

BNSDOCID: <WO 9842738A1>

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	AAGGCAAATT	CATGAAGGTG	CAAGTTTACC	GTTTTTTGAA	GTATTTGTTG	ATGCTCCTCT	540
5	GCATGTTTGT	GAACAGAGGG	ATGTCAAAGG	ACTCTACAAA	AAAGCCCGGG	CAGGAGAAAT	600
	TAAAGGTTTC	ACTGGGATCG	ATTCTGAATA	TGAAAAGCCA	GAGGCCCCTG	AGTTGGTGCT	660
	GAAAACAGAC	TCCTGTGATG	TAAATGACTG	TGTCCAGCAA	GTTGTGGAAC	TTCTACAGGA	720
10	ACGGGATATT	GTACCTGTGG	ATGCATCTTA	TGAAGTAAAA	GAACTATATG	TGCCAGAAAA	780
	TAAACTTCAT	TTGGCAAAAA	CAGATGCGGA	AACATTACCA	GCACTGAAAA	TTAATAAAGT	840
15	GGATATGCAG	TGGGTGCAGG	TTTTGGCAGA	AGGTTGGGCA	ACCCCATTGA	ATGGCTTTAT	900
	GAGAGAGAGG	GAGTACTTGC	AGTGCCTTCA	TTTTGATTGT	CTTCTGGATG	GAGGTGTCAT	960
	TAACTTGTCA	GTACCTATAG	TTCTGACTGC	GACTCATGAA	GATAAAGAGA	GGCTGGACGG	1020
20	CTGTACAGCA	TTTGCTCTGA	TGTATGAGGG	CCGCCGTGTG	GCCATTCTTC	GCAATCCAGA	1080
	GTTTTTTGAG	CACAGGAAAG	AGGAGCGCTG	TGCCAGACAG	TGGGGAACGA	CATGCAAGAA	1140
25	CCACCCCTAT	ATTAAGATGG	TGATGGAACA	AGGAGATTGG	CTGATTGGAG	GAGATCTTCA	1200
	AGTCTTGGAT	CGAGTTTATT	GGAATGATGG	TCTTGATCAG	TATCGTCTTA	CTCCTACTGA	1260
	GCTAAAGCAG	AAATTTAAAG	ATATGAATGC	TGATGCTGTC	TTTGCATTTC	AACTACGCAA	1320
30	CCCAGTGCAC	AATGGACATG	CCCTGTTAAT	GCAGGATACC	CATAAGCAAC	TTCTAGAGAG	1380
	GGGCTACCGG	CGCCCTGTCC	TCCTCCTCCA	CCCTCTGGGT	GGCTGGACAA	AGGATGACGA	1440
35	TGTTCCTTTG	ATGTGGCGTA	TGAAGCAGCA	TGCTGCAGTG	TTGGAGGAAG	GAGTTCTGAA	1500
	TCCTGAGACG	ACAGTGGTGG	CCATCTTCCC	ATCTCCCATG	ATGTATGCTG	GACCAACTGA	1560
	GGTCCAGTGG	CATTGCAGAG	CACGGATGGT	TGCAGGAGCC	AACTTTTACA	TTGTTGGACG	1620
40	AGACCCTGCT	GGCATGCCTC	ATCCAGAAAC	AGGGAAGGAT	CTTTATGAGC	CAAGTCATGG	1680
	TGCCAAAGTG	CTGACGATGG	CCCCTGGTTT	AATCACTTTG	GAAATAGTTC	CCTTTCGAGT	1740
45	TGCAGCTTAC	AACAAGAAAA	AGAAGCGTAT	GGACTACTAT	GACTCTGAAC	ACCATGAAGA	1800
	CTTTGAATTT	ATTTCAGGAA	CACGAATGCG	CAAACTTGCT	CGAGAAGGCC	AGAAACCACC	1860
	TGAAGGTTTC	ATGGCTCCCA	AGGCTTGGAC	CGTGCTGACA	GAATACTACA	AATCCTTGGA	1920
50	GAAAGCTTAG	GCTGTTAACC	CAGTCACTCC	ACCTTTGACA	CATTACTAGT	AACAAGAGGG	1980
	GACCACATAG	TCTCTGTTGG	CATTTCTTTG	TGGTGTCTGT	CTGGACATGC	TTCCTAAAAA	2040
55	CAGACCATTT	TCCTTAACTT	GCATCAGTTT	TGGTCTGCCT	TATGAGTTCT	GTTTTGAACA	2100
	AGTGTAACAC	ACTGATGGTT	TTAATGTATC	TTTTCCACTT	ATTATAGTTA	TATTCCTACA	2160
	ATACAATTTT	AAAATTGTCT	TTTTATATTA	TATTTATGCT	TCTGTGTCAT	GATTTTTTCA	2220
60	AGCTGTTATA	TTAGTTGTAA	CCAGTAGTAT	TCACATTAAA	TCTTGCTTTT	TTTCCCCTTA	2280

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10	AAAAAA						2527
	GTAAGCTCTG	AATGAACTTC	TTTACTCAAT	AAAATTAATT	TTTTGGCTTC	TTAAAAAAA	2520
5	CTTTCCAGTC	AGCTATTGGT	CTTTCCAGCT	GTTATAATCT	AAAGTATTCT	TATGATCTGT	2460
	AGACCTTTGT	AGCGATTAGA	TTTTTTTCT	ACATTGAAAA	TAGAAACTGC	TTCCTTTCTT	2400
	AAAAAAGAAA	AAAATTACCA	AACAATAAAC	TTGGCTAGAC	CTTGTTTGA	GGATTTTACA	2340

15 (2) INFORMATION FOR SEQ ID NO: 118:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1098 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 118:

CGCATCACAG ACAACCCAGA AGGAAAATGG TTGGGCAGAA CAGCAAGGGG TTCATATGGC 60 25 TATATTAAAA CAACTGCTGT AGAGATTNNC TATGATTCTT TGAAACTGAA AAAAGACTCT 120 CTTGGTGCCC CTTCAAGACC TATTGAAGAT GACCAAGAAG TATATGATGA TGTTGCAGAG 180 30 CAGGATGATA TTAGCAGCCA CAGTCAGAGT GGAAGTGGAG GGATATTCCC TCCACCACCA 240 GATGATGACA TTTATGATGG GATTGAAGAG GAAGATGCTG ATGATGGTTT CCCTGCTCCT 300 CCTAAACAAT TGGACATGGG AGATGAAGTT TACGATGATG TGGATACCTC TGATTTCCCT 35 360 GTTTCATCAG CAGAGATGAG TCAAGGAACT AATGTTGGAA AAGCTAAGAC AGAAGAAAAG 420 480 GACCTTAAGA AGCTAAAAAA GCAGRAAAAA GAARAAAAAG ACTTCAGGAA AAAATTTAAA 40 TATGATGGTG AAATTAGAGT CCTATATTCA ACTAAAGTTA CAACTTCCAT AACTTCTAAA AAGTGGGGAA CCAGAGATCT ACAGGTAAAA CCTGGTGAAT CTCTAGAAGT TATACAAACC 600 ACAGATGACA CAAAAGTTCT CTGCAGAAAT GAAGAAGGGA AATATGGTTA TGTCCTTCGG 660 45 AGTTACCTAG CGGACAATGA TGGAGAGATC TATGATGATA TTGCTGATGG CTGCATCTAT 720 GACAATGACT AGCACTCAAC TTTGGTCATT CTGCTGTGTT CATTAGGTGC CAATGTGAAG 780 50 TCTGGATTTT AATTGGCATG TTATTGGGTA TCMAGAAAAT TAATGCACAR AACCACTTAT 840 TATCATTTGT TATGAAATCC CAATTATCTT TACAAAGTGT TTAAAGTTTG AACATAGAAA 900 ATAATCTCTC TGCTTAATTG TTATCTCAGA AGACTACATT AGTGAGATGT AAGAATTATT 960 55 AAATATTCCA TTTCCGCTTT GGCTACAATT ATGAAGAAGT TGAAGGTACT TCTTTTAGAC 1020 1080

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GGGGGCCCGG TACCCAAT 1098

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(2) INFORMATION FOR SEQ ID NO: 119:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1679 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 119:

15 TCGACCCACG CGTCCGGCGA GATCCCTACC GCAGTAGCCG CCTCTGCCGC CGCGGAGCTT 60 CCCGAACCTC TTCAGCCGCC CGGAGCCGCT CCCGGAGCCC GGCCGTAGAG GCTGCAATCG 120 20 CAGCCGGGAG CCCGCAGCCC GCGCCCCGAG CCCGCCGCG CCCTTCGAGG GCGCCCCAGG 180 CCGCGCCATG GTGAAGGTGA CGTTCAACTC CGCTCTGGCC CAGAAGGAGG CCAAGAAGGA 240 CGAGCCCAAG AGCGGCGAGG AGGCGCTCAT CATCCCCCCC GACGCCGTCG CGGTGGACTG 300 25 CAAGGACCCA GATGATGTGG TACCAGTTGG CCAAAGAAGA GCCTGGTGTT GGTGCATGTG 360 CTTTGGACTA GCATTTATGC TTGCAGGTGT TATTCTAGGA GGAGCATACT TGTACAAATA 420 30 TTTTGCACTT CAACCAGATG ACGTGTACTA CTGTGGAATA AAGTACATCA AAGATGATGT 480 CATCTTAAAT GAGCCCTCTG CAGATGCCCC AGCTGCTCTC TACCAGACAA TTGAAGAAAA 540 TATTAAAATC TTTGAAGAAG AAGAAGTTGA ATTTATCAGT GTGCCTGTCC CAGAGTTTGC 600 35 AGATAGTGAT CCTGCCAACA TTGTTCATGA CTTTAACAAG AAACTTACAG CCTATTTAGA 660 TCTTAACCTG GATAAGTGCT ATGTGATCCC TCTGAACACT TCCATTGTTA TGCCACCCAG 720 40 AAACCTACTG GAGTTACTTA TTAACATCAA GGCTGGAACC TATTTGCCTC AGTCCTATCT 780 GATTCATGAG CACATGGTTA TTACTGATCG CATTGAAAAC ATTGATCACC TGGGTTTCTT 840 TATTTATCGA CTGTGTCATG ACAAGGAAAC TTACAAACTG CAACGCAGAG AAACTATTAA 900 45 AGGTATTCAG AAACGTGAAG CCAGCAATTG TTTCGCAATT CGGCATTTTG AAAACAAATT 960 TGCCGTGGAA ACTTTAATTT GTTCTTGAAC AGTCAAGAAA AACATTATTG AGGAAAATTA 1020 50 ATATCACAGC ATAACCCCAC CCTTTACATT TTGTGCAGTG ATTATTTTTT AAAGTCTTCT 1080 TTCATGTAAG TAGCAAACAG GGCTTTACTA TCTTTTCATC TCATTAATTC AATTAAAACC 1140 ATTACCTTAA AATTITITITC TITCGAAGTG TGGTGTCTTT TATATTIGAA TTAGTAACTG 1200 55 TATGAAGTCA TAGATAATAG TACATGTCAC CTTAGGTAGT AGGAAGAATT ACAATTTCTT 1260 TAAATCATTT ATCTGGATTT TTATGTTTTA TTAGCATTTT CAAGAAGACG GATTATCTAG 1320 60 AGAATAATCA TATATATGCA TACGTAAAAA TGGACCACAG TGACTTATTT GTAGTTGTTA 1380

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	GTTGCCCTGC	TACCTAGTTT	GTTAGTGCAT	TTGAGCACAC	ATTTTAATTT	TCCTCTAATT	1440
ہے	AAAATGTGCA	GTATTTTCAG	TGTCAAATAT	ATTTAACTAT	TTAGAGAATG	ATTTCCACCT	1500
3	TTATGTTTTA	ATATCCTAGG	CATCTGCTGT	AATAATATT	TAGAAAATGT	TTGGAATTTA	1560
	AGAAATAACT	TGTGTTACTA	ATTTGTATAA	CCCATATCTG	TGCAATGGAA	TATAAATATC	1620
10	ACAAAGTTGT	TTAAMWAAAA	AAAAAAAA	AAAAAAAA	AAAAAAAA	AAAAAAAN	1679

15 (2) INFORMATION FOR SEQ ID NO: 120:

20

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1308 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 120:

25	TTGGCANCNG	GGAGAGGGAA	AGAGGAGGAA	ATGGGGTTTG	AGGACCATGG	CTTACCTTTC	60
	CTGCCTTTGA	CCCATCACAC	CCCATTTCCT	CCTCTTTCCC	TCTCCCCGCT	GCCAAAAAA	120
20	AAAAAAAAGG	AAACGTTTAT	CATGAATCAA	CAGGGTTTCA	GTCCTTATCA	AAGAGAGATG	180
30	TGGAAAGAGC	TAAAGAAACC	ACCCTTTGTT	CCCAACTCCA	CTTTACCCAT	ATTTTATGCA	240
	ACACAAACAC	TGTCCTTTTG	GGTCCCTTTC	TTACAGATGG	ACCTCTTGAG	AAGAATTATC	300
35	GTATTCCACG	TTTTTAGCCC	TCAGGTTACC	AAGATAAATA	TATGTATATA	TAACCTTTAT	360
	TATTGCTATA	TCTTTGTGGA	TAATACATTC	AGGTGGTGCT	GGGTGATTTA	TTATAATCTG	420
40	AACCTAGGTA	TATCCTTTGG	TCTTCCACAG	TCATGTTGAG	GTGGGCTCCC	TGGTATGGTA	480
40	AAAAGCCAGG	TATAATGTAA	CTTCACCCCA	GCCTTTGTAC	TAAGCTCTTG	ATAGTGGATA	540
	TACTCTTTTA	AGTTTAGCCC	CAATATAGGG	TAATGGAAAT	TTCCTGCCCT	CTGGGTTCCC	600
45	CATTTTTACT	' ATTAAGAAGA	CCAGTGATAA	TTTAATAATG	CCACCAACTC	TGGCTTAGTT	660
	AAGTGAGAGT	GTGAACTGTG	TGGCAAGAGA	GCCTCACACC	TCACTAGGTG	CAGAGAGCCC	720
50	AGGCCTTATO	TTAAAATCAT	GCACTTGAAA	AGCAAACCTT	AATCTGCAAA	GACAGCAGCA	780
30	AGCATTATAC	GGTCATCTTG	AATGATCCCT	TTGAAATTT	TITITITGTT	GTTTGTTTAA	840
	ATCAAGCCTC	G AGGCTGGTGA	ACAGTAGCTA	A CACACCCATA	TTGTGTGTTC	TGTGAATGCT	900
55	AGCTCTCTTC	AATTTGGATA	TTGGTTATT	TTTATAGAGT	GTAAACCAAG	TTTTATATTC	960
	TGCAATGCG	A ACAGGTACCT	ATCTGTTTC	CAAATAAAA 1	GTTTACATTC	ATTATGGGGT	1020
60	ATGTATGAC	C TTCATTTCC	C AAGAAATAGA	A ACTCTAGCT	AGAATTATGO	ATGCTCTAAA	1080

	ATGTCAGAAT GGGAACTCTC CTCGAAGTTC TCCCAAACTC AGAGACAGCA CTGCCTTCTC	1140
	CTAAATGATT ATTCTTTTCT CCCTGTTTTC TGGTATTTTC TAGGCATCCT TCTCACCACA	1200
5	GCCATAACCC TITITTACTT CCATTAGGCC GTATAACTGG NGGGACNGCT GGTCGGTATA	1260
	TAATACTGGT WCCAACAMAG GGGTTCTGGA TGTACACMAG GTTATCTT	1308
10		
10		
	(2) INFORMATION FOR SEQ ID NO: 121:	
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 1411 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 121:	
	GGCACAGGAG CGACCCGGGA GAAGGAGGGC CAMGAKGCGG AAGCGGAGGA GTCTCCAGGA	60
25	GACCCGGGGA CAGCATCGCC CAGGCCCCTG TTTGCAGGCC TTTCAGATAT ATCCATCTCA	120
	CAAGACATCC CCGTAGAAGG AGAAATCACC ATTCCTATGA GATCTCGCAT CCGGGAGTTT	180
	GACAGCTCCA CATTAAATGA ATCTGTTCGC AATACCATCA TGCGTGATCT AAAAGCTGTT	240
30	GGGAAAAAT TCATGCATGT TTTGTACCCA AGGAAAAGTA ATACTCTTTT GAGAGATTGG	300
	GATTTGTGGG GCCCTTTGAT CCTTTGTGTG ACACTCGCAT TAATGCTGCA AAGAGACTCT	360
35	GCAGATAGTG AAAAAGATGG AGGGCCCCAA TTTGCAGAGG TGTTTGTCAT TGTCTGGTTT	420
	GGTGCAGTTA CCATCACCCT CAACTCAAAA CTTCTTGGAG GGAACATATC TTTTTTCAG	480
	AGCCTCTGTG TGCTGGGTTA CTGTATACTT CCCTTGACAG TAGCAATGCT GATTTGCCGG	540
40	CTGGTACTTT TGGCTGATCC AGGACCTGTA AACTTCATGG TTCGGCTTTT TGTGGTGATT	600
	GTGATGTTTG CCTGGTCTAT AGTTGCCTCC ACAGCTTTCC TTGCTGATAG CCAGCCTCCA	660
45	AACCGCAGAG CCCTAGCTGT TTATCCTGTT TTCCTGTTTT ACTTTGTCAT CAGTTGGATG	720
	ATTCTCACCT TTACTCCTCA GTAAATCAGG AATGGGAAAT TAAAAACCAG TGAATTGAAA	780
	GCACATCTGA AAGATGCAAT TCACCATGGA GCTTTGTCTC TGGCCCTTAT TTGTCTAATT	840
50	TTGGAGGTAT TTGATAACTG AGTAGGTGAG GAGATTAAAA GGGAGCCATA TAGCACTGTC	900
	ACCCCTTATT TGAGGAACTG ATGTTTGAAA GGCTGTTCTT TTCTCTCTTA ATGTCATTTC	960
55	TTTAAAAATA CATGTGCATA CTACACACAG TATATAATGC CTCCTTAAGG CATGATGGAG	1020
	TCACCGTGGT CCATTTGGGT GACAACCAGT GACTTGGGAA GCACATAGAT ACATCTTACA	1080
	AGTTGAATAG AGTTGATAAC TATTTTCAGT TTTGAGAATA CCAGTTCAGG TGCAGCTCTT	1140
60	AAACACATTG CCTTATGACT ATTAGAATAT GCCTCTCTTT TCATAAATAA AAATACATGG	1200

	TCTATATCCA TTTTCTTTTA TTTCTCTCTC TTAAGCTTAA AAAGGCAATG AGAGAGGTTA	1260
-	GGAGTGGGTT CATACACGGA GAATGAGAAA ACATGCATTA ACCAATATTC AGATTTTGAT	1320
5	CAGGGGAAAT TCTAYACTTG TTGCAAAAAA AAAAAAAAAA AAACTCGAGG GGGGCCCGGT	1380
	ACCCAATCGC NGTATATGAT CGNAAACAAT C	1411
10		
	(2) INFORMATION FOR SEQ ID NO: 122:	
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 2256 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 122:	
	GCTTTGGCTT TTTTTGGCGG ACTGGGGCGC CCTCCGGAAG CGTTTCCAAC TTTCCAGAAG	60
25	TTTCTCGGGA CGGGCAGGAG GGGGTGGGGA CTGCCATATA TAGATCCCGG GAGCAGGGGA	120
	GCGGGCTAAG AGTAGAATCG TGTCGCGGCT CGAGAGCGAG AGTCACGTCC CGGCGCTAGC	180
20	CAGCCCGACC CAGGCCCACC GTGGTGCACG CAAACCACTT CCTGGCCATG CGCTCCCTCC	240
30	TGCTTCTCAG CGCCTTCTGC CTCCTGGAGG CGGCCCTGGC CGCCGAGGTG AAGAAACCTG	300
	CAGCCGCAGC AGCTCCTGGC ACTGCGGAGA AGTTGAGCCC CAAGGCGGCC ACGCTTGCCG	360
35	AGCGCANGCC GGCCTGGCCT TCAGCTTGTA CCAGGCCATG GCCAAGGACC AGGCAGTGGA	420
	GAACATCCTG GTGTCACCCG TGGTGGTGGC CTCGTCGCTG GGGCTCGTGT CGCTGGGCGG	480
40	CAAGGCGACC ACGGCGTCGC AGGCCAAGGC AGTGCTGAGC GCCGAGCAGC TGCGCGACGA	540
40	GGAGGTGCAC GCCGGCCTGG GCGAGCTGCT GCGCTCACTC AGCAACTCGA CGGCGCGCAA	600
	CGTGACCTGG AAGCTGGGCA GCCGACTGTA CGGACCCAGC TCAGTGAGCT TCGCTGATGA	660
45	CTTCGTGCGC ACAGCAAGCA GCACTACAAC TGCGAGCACT CCAAGATCAA CTTCCGCGAC	720
	AAGCGCAGNG CGCTGCAGTC CATCAACGAG TGGGCCGCGC AGACCACCGA CGGCAAGCTG	780
50	CCCGAGGTCA CCAAGGACGT GGAGCGCACG GACGGCGCCC TGCTAGTCAA CGCCATGTTC	840
50	TTCAAGCCAC ACTGGGATGA GAAATTCCAC CACAAGATGG TGGACAACCG TGGCTTCATG	900
	GTGACTCGGT CCTATACYGT GGGTGTCATG ATGATGCACC GGACAGGCCT CTACAACTAC	960
55	TACGACGACG AGAAGGAAAA GCTGCAAATC GTGGAGATGC CCCTGGCCCA CAAGCTCTCC	1020
	AGCCTCATCA TCCTCATGCC CCATCACGTG GAGCCTCTCG AGCGCCTTGA AAAGCTGCTA	1080
60	ACCAAAGAGC AGCTGAAGAT CTGGATGGGG AAGATGCAGA AGAAGGCTGT TGCCATCTCC	1140

	TTGCCCAAGG	GTGTGGTGGA	GGTGACCCAT	GACCTGCAGA	AACACCTGGC	TGGGCTGGGC	1200
	CTGACTGAGG	CCATTGACAA	GAACAAGGCC	GACTTRTCAC	GCATGTCAGG	CAAGAAGGAC	1260
5	CTGTACCTGG	CCAGCGTGTT	CCACGCCACC	GCCTTTGAGT	TGGACACAGA	TGGCAACCCC	1320
	TTTGACCAGG	ACATCTACGG	GCGCGAGGAG	CTGCGCANCC	CAAGCTGTTC	TACGCCGACC	1380
10	ACCCCTTCAT	CTTCCTAGTG	CGGGACACCC	AAAGCGGCTC	CCTGCTATTC	ATTGGGCGCC	1440
	TGGTCCGGCC	TAAGGGTGAC	AAGATGCGAG	ACGAGTTATA	GGGCCTCAGG	GTGCACACAG	1500
	GATGGCAGGA	GGCATCCAAA	GGCTCCTGAG	ACACATGGGT	GCTATTGGGG	TTGGGGGGGA	1560
15	GGTGAGÇTAC	CAGCCTTGGA	TACTCCATGG	GGTGGGGGTG	GAAAARCAGA	CCGGGGTTCC	1620
	CGTGTGCCTG	AGCGGACCTT	CCCAGCTAGA	ATTCACTCCA	CTTGGACATG	GGCCCCAGAT	1680
20	ACCATGATGC	TGAGCCCGGA	AACTCCACAT	CCTGTGGGAC	CTGGGCCATA	GTCATTCTGC	1740
	CTGCCCTGAA	AGTCCCAGAT	CAAGCCTGCC	TCAATCAGTA	TTCATATTTA	TAGCCAGGTA	1800
	CCTTCTCACC	TGTGAGACCA	AATTGAGCTA	GGGGGGTCAG	CCAGCCCTCT	TCTGACACTA	1860
25	AAACACCTCA	GCTGCCTCCC	CAGCTCTATC	CCAACCTCTC	CCAACTATAA	AACTAGGTGC	1920
	TGCAGCCCCT	GGGACCAGGC	ACCCCCAGAA	TGACCTGGCC	GCAGTGAGGC	GGATTGAGAA	1980
30	GGAGCTCCCA	GGAGGGGCTT	CTGGGCAGAC	TCTGGTCAAG	AAGCATCGTG	TCTGGCGTTG	2040
	TGGGGATGAA	CTTTTTGTTT	TGTTTCTTCC	TTTTTTAGTT	CTTCAAAGAT	AGGGAGGGAA	2100
	GGGGGAACAT	GAGCCTTTGT	TGCTATCAAT	CCAAGAACTT	ATTTGTACAT	TTTTTTTC	2160
35	AATAAAACTT	TTCCAATGAC	AAAAAAAA	AAAAAAAAA	AAAAAGGGGS	GGCCGCTCC	2220
	TAGAGGGATC	CCTCCGANGG	NGCCCAATCG	AAAATN			2256
40							
	(2) INFORMA	TION FOR SE	O ID NO: 12	3:			
45		SEQUENCE CH (A) LENC (B) TYPE (C) STRA		CCS: se pairs acid double			
50	(xi)	SEQUENCE D	ESCRIPTION:	SEQ ID NO:	123:		
	ATGCGCTCCC	TCCTGCTTCT	CAGCGCCTTC	TGCCTCCTGG	AGGCGGCCCT	GGCCGCCGAG	60
55	GTGAAGAAAC	CTGCAGCCGC	AGCAGCTCCT	GGCACTGCGG	AGAAGTTGAG	CCCCAAGGCG	120
	GCCACGCTTG	CCGAGCGCAA	GCGGCCTGGC	CTTCAGCTTG	TACCAGGCCA	TGGCCAAGGA	180
	CCAGGCAGTG	GAGAACATCC	TGGTGTCACC	CGTGGTGGTG	GCCTCGTCGC	TGGGGCTCGT	240
60	GTCGCTGGGC	GGCAAGGCGA	CCACGGCGTC	GCAGGCCAAG	GCAGTGCTGA	GCGCCGAGCA	300

	GCTGCGCGAC GAGGAGGTGC ACGCCGGCCT GGGCGAGCTG CTGCGCTCAC TCAGCAACTC	360
5	CACGGCGCGC AACGTGACCT GGAAGCTGGG CAGCCGACTG TACGGACCCA GCTCAGTGAG	420
3	CTTCGCTGAT GACTTCGTGC GCAGCAGCAA GCAGCACTAC AACTGCGAGC ACTCCAAGAT	480
	CAACTTCCGC GACAAGCGCA GCGCGCTGCA GTCCATCAAC GAGTGGGCCG CGCAGACCAC	540
10	CGACGCCAAG CTGCCCGAGG TCACCAAGGA CGTGGAGCGC ACGGACGGCG CCCTGTTAGT	600
	CAACGCCATG TTCTTCAAGC CACACTGGGA TGAGAAATTC CACCACAAGA TGGTGGACAA	660
15	CCGTGGCTTC ATGGTGACTC GGTCCTATAC CGTGGGTGTC ATGATGATGC ACCGGACAGG	720
13	CCTCTACAAC TACTACGACG ACGAGAAGGA AAAGCTGCAA ATCGTGGAGA TGCCCCTGGC	780
	CCACAAGCTC TCCAGCCTCA TCATCCTCAT GCCCCATCAC GTGGAGCCT	829
20		
	(2) INFORMATION FOR SEQ ID NO: 124:	
25	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 2223 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
30		
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 124:	
	(XI) DECOMICE DESCRIPTION DEC ES TOTAL	
	CCTCCGGAAG CGTTTCCAAC TTTCCAGAAG TTTCTCGGGA CGGGCAGGAG GGGTGGGGA	60
35		60 120
35	CCTCCGGAAG CGTTTCCAAC TTTCCAGAAG TTTCTCGGGA CGGGCAGGAG GGGGTGGGGA	
	CCTCCGGAAG CGTTTCCAAC TTTCCAGAAG TTTCTCGGGA CGGGCAGGAG GGGGTGGGGA CTGCCATATA TAGATCCCGG GAGCAGGGGA GCGGGCTAAG AGTAGAATCG TGTCGCGGCT	120
35 40	CCTCCGGAAG CGTTTCCAAC TTTCCAGAAG TTTCTCGGGA CGGGCAGGAG GGGGTGGGGA CTGCCATATA TAGATCCCGG GAGCAGGGGA GCGGGCTAAG AGTAGAATCG TGTCGCGGCT CGAGAGCGAG AGTCACGTCC CGGCGCTAGC CAGCCCGACC CAGGCCCACC GTGGTGCACG	120 180
	CCTCCGGAAG CGTTTCCAAC TTTCCAGAAG TTTCTCGGGA CGGGCAGGAG GGGGTGGGGA CTGCCATATA TAGATCCCGG GAGCAGGGGA GCGGGCTAAG AGTAGAATCG TGTCGCGGCT CGAGAGCGAG AGTCACGTCC CGGCGCTAGC CAGCCCGACC CAGGCCCACC GTGGTGCACG CAAACCACTT CCTGGCCATG CGCTCCCTCC TGCTTCTCAG CGCCTTCTGC CTCCTGGAGG	120 180 240
	CCTCCGGAAG CGTTTCCAAC TTTCCAGAAG TTTCTCGGGA CGGGCAGGAG GGGGTGGGGA CTGCCATATA TAGATCCCGG GAGCAGGGGA GCGGGCTAAG AGTAGAATCG TGTCGCGGCT CGAGAGCGAG AGTCACGTCC CGGCGCTAGC CAGCCCGACC CAGGCCCACC GTGGTGCACG CAAACCACTT CCTGGCCATG CGCTCCCTCC TGCTTCTCAG CGCCTTCTGC CTCCTGGAGG CGGCCCTGGC CGCCGAGGTG AAGAAACCTG CAGCCGCAGC AGCTCCTGGC ACTGCGGAGA	120 180 240 300
40	CCTCCGGAAG CGTTTCCAAC TTTCCAGAAG TTTCTCGGGA CGGGCAGGAG GGGGTGGGGA CTGCCATATA TAGATCCCGG GAGCAGGGGA GCGGGCTAAG AGTAGAATCG TGTCGCGGCT CGAGAGCGAG AGTCACGTCC CGGCGCTAGC CAGCCCGACC CAGGCCCACC GTGGTGCACG CAAACCACTT CCTGGCCATG CGCTCCCTCC TGCTTCTCAG CGCCTTCTGC CTCCTGGAGG CGGCCCTGGC CGCCGAGGTG AAGAAACCTG CAGCCGCAGC AGCTCCTGGC ACTGCGGAGA AGTTGAGCCC CAAGGCGGCC ACGCTTGCCG AGCGCAGNCG GCCTGGCCTT CAGCTTGTAC	120 180 240 300 360
40 45	CCTCCGGAAG CGTTTCCAAC TTTCCAGAAG TTTCTCGGGA CGGGCAGGAG GGGGTGGGGA CTGCCATATA TAGATCCCGG GAGCAGGGGA GCGGGCTAAG AGTAGAATCG TGTCGCGGCT CGAGAGCGAG AGTCACGTCC CGGCGCTAGC CAGCCCGACC CAGGCCCACC GTGGTGCACG CAAACCACTT CCTGGCCATG CGCTCCCTCC TGCTTCTCAG CGCCTTCTGC CTCCTGGAGG CGGCCCTGGC CGCCGAGGTG AAGAAACCTG CAGCCGCAGC AGCTCCTGGC ACTGCGGAGA AGTTGAGCCC CAAGGCCGCC ACGCTTGCCG AGCGCAGNCG GCCTGGCCTT CAGCTTGTAC CAGGCCATGG CCAAGGACCA GGCAGTGGAG AACATCCTGG TGTCACCCGT GGTGGTGGCC	120 180 240 300 360 420
40	CCTCCGGAAG CGTTTCCAAC TTTCCAGAAG TTTCTCGGGA CGGGCAGGAG GGGGTGGGGA CTGCCATATA TAGATCCCGG GAGCAGGGGA GCGGGCTAAG AGTAGAATCG TGTCGCGGCT CGAGAGCGAG AGTCACGTCC CGGCGCTAGC CAGCCCGACC CAGGCCCACC GTGGTGCACG CAAACCACTT CCTGGCCATG CGCTCCCTCC TGCTTCTCAG CGCCTTCTGC CTCCTGGAGG CGGCCCTGGC CGCCGAGGTG AAGAAACCTG CAGCCGCAGC AGCTCCTGGC ACTGCGGAGA AGTTGAGCCC CAAGGCGGCC ACGCTTGCCG AGCGCAGNCG GCCTGGCCTT CAGCTTGTAC CAGGCCATGG CCAAGGACCA GGCAGTGGAG AACATCCTGG TGTCACCCGT GGTGGTGGCC TCGTCGCTGG GGCTCGTGTC GCTGGGCGGC AAGGCGACCA CGGCGTCGCA GGCCAAGGCA	120 180 240 300 360 420 480
40 45	CCTCCGGAAG CGTTTCCAAC TTTCCAGAAG TTTCTCGGGA CGGGCAGGAG GGGGTGGGGA CTGCCATATA TAGATCCCGG GAGCAGGGGA GCGGGCTAAG AGTAGAATCG TGTCGCGGCT CGAGAGCGAG AGTCACGTCC CGGCGCTAGC CAGCCCGACC CAGGCCCACC GTGGTGCACG CAAACCACTT CCTGGCCATG CGCTCCCTCC TGCTTCTCAG CGCCTTCTGC CTCCTGGAGG CGGCCCTGGC CGCCGAGGTG AAGAAACCTG CAGCCGCAGC AGCTCCTGGC ACTGCGGAGA AGTTGAGCCC CAAGGCGGCC ACGCTTGCCG AGCGCAGNCG GCCTGGCCTT CAGCTTGTAC CAGGCCATGG CCAAGGACCA GGCAGTGGAG AACATCCTGG TGTCACCCGT GGTGGTGGCC TCGTCGCTGG GGCTCGTGTC GCTGGGCGGC AAGGCGACCA CGGCGTCGCA GGCCAAGGCA GTGCTGAGCG CCGAGCAGCT GCGCGACGAG GAGGTGCACG CCGGCCTGGG CGAGCTGCTG	120 180 240 300 360 420 480 540
40 45	CCTCCGGAAG CGTTTCCAAC TTTCCAGAAG TTTCTCGGGA CGGGCAGGAG GGGGTGGGGA CTGCCATATA TAGATCCCGG GAGCAGGGGA GCGGGCTAAG AGTAGAATCG TGTCGCGGCT CGAGAGCGAG AGTCACGTCC CGGCGCTAGC CAGCCCGACC CAGGCCCACC GTGGTGCACG CAAACCACTT CCTGGCCATG CGCTCCCTCC TGCTTCTCAG CGCCTTCTGC CTCCTGGAGG CGGCCCTGGC CGCCGAGGTG AAGAAACCTG CAGCCGCAGC AGCTCCTGGC ACTGCGGAGA AGTTGAGCCC CAAGGCGGCC ACGCTTGCCG AGCGCAGAC GCCTGGCCTT CAGCTTGTAC CAGGCCATGG CCAAGGACCA GGCAGTGGAG AACATCCTGG TGTCACCCGT GGTGGTGGCC TCGTCGCTGG GGCTCGTGTC GCTGGGCGGC AAGGCGACCA CGGCGTCGCA GGCCAAGGCA GTGCTGAGCG CCGAGCAGCT GCGCGACGAG GAGGTGCACG CCGGCCTGGG CGAGCTGCTG CGCTCACTCA GCAACTCSAC GGCGCGCAAC GTGACCTGGA AGCTGGGCAG CCGACTGTAC	120 180 240 300 360 420 480 540

CGGCGCCCTG YTAGTCAACG CCATGTTCTT CAAGCCACAC TGGGATGAGA AATTCCACCA

	CAAGATGGTG	GACAACCGTG	GCTTCATGGT	GACTCGGTCC	TATACYGTGG	GTGTCATGAT	900
	GATGCACCGG	ACAGGCCTCT	ACAACTACTA	CGACGACGAG	AAGGAAAAGC	TGCAAATCGT	960
5	GGAGATGCCC	CTGGCCCACA	AGCTCTCCAG	CCTCATCATC	CTCATGCCCC	ATCACGTGGA	1020
	GCCTCTCGAG	CGCCTTGAAA	AGCTGCTAAC	CAAAGAGCAG	CTGAAGATCT	GGATGGGGAA	1080
10	GATGCAGAAG	AAGGCTGTTG	CCATCTCCTT	GCCCAAGGGT	GTGGTGGAGG	TGACCCATGA	1140
10	CCTGCAGAAA	CACCTGGCTG	GGCTGGGCCT	GACTGAGGCC	ATTGACAAGA	ACAAGGCCGA	1200
	CTTRTCACGC	ATGTCAGGCA	AGAAGGACCT	GTACCTGGCC	AGCGTGTTCC	ACGCCACCGC	1260
15	CTTTGAGTTG	GACACAGATG	GCAACCCCTT	TGACCAGGAC	ATCTACGGGC	GCGAGGAGCT	1320
	GCGCASCCCA	AGCTGTTCTA	CGCCGACCAC	CCCTTCATCT	TCCTAGTGCG	GGACACCCAA	1380
20	AGCGGCTCCC	TGCTATTCAT	TGGGCGCCTG	GTCCGGCCTA	AGGGTGACAA	GATGCGAGAC	1440
20	GAGTTATAGG	GCCTCAGGGT	GCACACAGGA	TGGCAGGAGG	CATCCAAAGG	CTCCTGAGAC	1500
	ACATGGGTGC	TATTGGGGTT	GGGGGGAGG	TGAGGTACCA	GCCTTGGATA	CTCCATGGGG	1560
25	TGGGGTGGA	AAARCAGACC	GGGGTTCCCG	TGTGCCTGAG	CGGACCTTCC	CAGCTAGAAT	1620
	TCACTCCACT	TGGACATGGG	CCCCAGATAC	CATGATGCTG	AGCCCGGAAA	CTCCACATCC	1680
30	TGTGGGACCT	GGGCCATAGT	CATTCTGCCT	GCCCTGAAAG	TCCCAGATCA	AGCCTGCCTC	1740
	AATCAGTATT	CATATTTATA	GCCAGGTACC	TTCTCACCTG	TGAGACCAAA	TTGAGCTAGG	1800
	GGGGTCAGCC	AGCCCTCTTC	TGACACTAAA	ACACCTCAGC	TGCCTCCCCA	GCTCTATCCC	1860
35	AACCTCTCCC	AACTATAAAA	CTAGGTGCTG	CAGCCCCTGG	GACCAGGCAC	CCCCAGAATG	1920
	ACCTGGCCGC	AGTGAGGCGG	ATTGAGAAGG	AGCTCCCAGG	AGGGGCTTCT	GGGCAGACTC	1980
40	TGGTCAAGAA	GCATCGTGTC	TGGCGTTGTG	GGGATGAACT	TTTTGTTTTG	TTTCTTCCTT	2040
	TTTTAGTTCT	TCAAAGATAG	GGAGGGAAGG	GGGAACATGA	GCCTTTGTTG	CTATCAATCC	2100
	AAGAACTTAT	TTGTACATTT	TTTTTTCAA	TAAAACTTTT	CCAATGACAA	AAAAAAAA	2160
45	АААААААА	MWMGGGGSGG	GCCGCTCCTA	GAGGGATCCC	TCCGANGGNG	CCCAATCGAA	2220
	AAT						2223

55

(2) INFORMATION FOR SEQ ID NO: 125:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 31 amino acids

(B) TYPE: amino acid

(D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 125:

 $60\,$ Met Lys Lys Gln Ser Lys Arg Cys Leu Trp Lys Pro Pro Gly Ser Leu

	1		!	5				10					15	
5	Arg Arg	Leu 1	Trp Trj 20	p Met	Arg	Ala	Leu 25	Leu	Ile	Leu	Lys	Tyr 30	Ile	
	(2) INF	ORMAT	ION FO	R SEQ	ID N	ю: 1	.26:							
10			(B)	LENGT TYPE: TOPOL	H: 4 ami OGY:	5 am no a lin	ino a cid ear	acid		: 120	6:			
15	Met Lys		Ser Le									Met	Leu 15	Leu
20	His Lev	ı Thr	Ala Al 20	a Phe	Leu	Gln	Arg 25	Ala	His	Xaa	Ile	Leu 30	Thr	Thr
	Arg Met	Ser : 35	Leu Gl	y Phe	Gln	Ser 40	Pro	His	Leu	Thr	Met 45			
25	(2) INF	FORMAT	'ION FO	OR SEQ	ID I	NO:	127:							
30			(B)	LENGT TYPE: TOPOI	TH: 3 ami LOGY:	9 am no a lin	nino Icid Iear	acid): 12	:7 :			
35	Met Hi:	s Asn	Gln Aı	rg Gln 5	Val	Phe	Leu	Phe 10		Leu	Phe	Ser	Asn 15	
40	Leu Le		20				Gly 25		Leu	Leu	ı Ala	. Ala 30		Tyr
45	(2) IN	FORMAT	rion F	OR SE(Q ID	NO:	128:							
50			(B)	LENG TYPE TOPO	TH: : am LOGY	23 au ino a : li:	mino acid near	aci		O: 1	28:			
55	Met Ar 1	g Lys	Lys P	he Le	u Lei	ı Ala	a Glr	n Vai		e Lei	u Sei	r Lei	ı Ser 15	
	Met Pr	ro Ser	Met P 20	ro Va	l Thi	r								
60														

	(2)	INF	ORMA	TION	FOR	SEQ	ID :	NO:	129:							
5				(A) L B) T D) T	ENGT YPE : OPOL	H: 1 ami OGY:	.10 a no a lin	mino .cid .ear	aci						
			(xi)			E DE										
10	Met 1	Val	Leu	Leu	Суs 5	Leu	Leu	Leu	Val	Pro 10	Leu	Leu	Leu	Ser	Leu 15	Ph
15	Val	Leu	Gly	Leu 20	Phe	Leu	Trp	Phe	Leu 25	Lys	Arg	Glu	Arg	Gln 30	Glu	Gl
	Tyr	Ile	Glu 35	Glu	Lys	Lys	Arg	Val 40	Asp	Ile	Cys	Arg	Glu 45	Thr	Pro	As
20	Ile	Cys 50	Pro	His	Ser	Gly	Glu 55	Asn	Thr	Glu	Tyr	Asp 60	Thr	Ile	Pro	Hi
	Thr 65	Asn	Arg	Thr	Ile	Leu 70	Lys	Glu	Asp	Pro	Ala 75	Asn	Thr	Val	Tyr	Se:
25	Thr	Val	Glu	Ile	Pro 85	Lys	Lys	Met	Glu	Asn 90	Pro	His	Ser	Leu	Leu 95	Th
30	Met	Pro	Asp	Thr 100	Pro	Arg	Leu	Phe	Ala 105	Tyr	Glu	Asn	Val	Ile 110		
	(2)	INF	ORMA!	rion	FOR	SEQ	ID 1	NO: 1	L30:							
35			(i)	(. (:	A) L B) T	CHAI ENGT YPE:	H: 6 ami	3 am no a	ino . cid		s					
40			(xi)							EQ II	ОИС	: 13	0:			
	Met 1	Leu	Leu	Leu	Phe 5	Ile	Tyr	Phe	Tyr	Ser 10	His	Pro	Ala	Pro	Val 15	Pro
45	Ala	Gly	Ala	Thr 20		Lys	Pro	Arg	Туr 25	Arg	Val	Ile	Thr	Cys 30	Gly	Pro
	Ala	Ser	Val 35	Phe	Ser	Thr	Ser	Phe 40	Ser	His	Ser	Pro	Pro 45	Ala	Arg	Cys
50	Leu	Gly 50	Arg	Leu	Glu	Gln	Met 55	Phe	His	Phe	Gly	Leu 60	Ala	Ser	Gly	
55	(2)	INFO	ORMAT	rion	FOR	SEQ	ID N	JO: 1	.31:							
			(i) :			CHAI ENGT					S					
50				(1	в) Т	YPE:	ami	no a	cid							

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 131: Met Pro Phe Pro Ile Ser Ile Leu Gln Leu Cys Leu Gln Ile Ser Asn 10 5 Leu Ser Phe Cys Leu Gln Lys Ile Tyr Lys Ile Pro Phe Val 25 10 (2) INFORMATION FOR SEQ ID NO: 132: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 53 amino acids (B) TYPE: amino acid 15 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 132: Met Ala Ala Ala Cys Arg Ser Val Lys Gly Leu Val Ala Val Ile Thr 20 1 Gly Gly Ala Ser Gly Leu Gly Leu Ala Thr Ala Asp Asp Leu Trp Gly Arg Glu Pro Leu Cys Phe Trp Thr Cys Pro Thr Arg Val Gly Arg 25 40 Pro Lys Pro Arg Ser 50 30 (2) INFORMATION FOR SEQ ID NO: 133: (i) SEQUENCE CHARACTERISTICS: 35 (A) LENGTH: 57 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 133: 40 Met Leu Leu Val Tyr Asp Leu Tyr Leu Xaa Pro Lys Leu Trp Ala Leu Ala Thr Pro Gln Lys Asn Gly Lys Gly Ala Arg Xaa Gly Asp Gly Thr 45 25 Pro Ala Gln Ala Phe Trp Asp Phe Trp Ser His Leu Ile Ser Ala Asp 40 Pro Gln Thr Trp Glu Arg Ala Ala Pro 50 50 (2) INFORMATION FOR SEQ ID NO: 134: 55 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 216 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear 60

			(xi)	SEQ	UENC	E DE	SCRI	PTIO	N: S	EQ I	D NO	: 13	4:			
5	Met 1	Arg	Leu	Ser	Ala 5	Leu	Leu	Ala	Leu	Ala 10	Ser	Lys	Val	Thr	Leu 15	Pro
٥	Pro	His	Tyr	Arg 20	Tyr	Gly	Met	Ser	Pro 25	Pro	Gly	Ser	Val	Ala 30	Asp	Lys
10	Arg	Lys	Asn 35	Pro	Pro	Trp	Ile	Arg 40	Arg	Arg	Pro	Val	Val 45	Val	Glu	Pro
	Ile	Ser 50	Asp	Glu	Asp	Trp	Tyr 55	Leu	Phe	Cys	Gly	Asp 60	Thr	Val	Glu	Ile
15	Leu 65	Glu	Gly	Lys	Asp	Ala 70	Gly	Lys	Gln	Gly	Lys 75	Val	Val	Gln	Val	Il€ 80
20	Arg	Gln	Arg	Asn	Trp 85	Val	Val	Val	Gly	Gly 90	Leu	Asn	Thr	His	Tyr 95	Arg
	Tyr	Ile	Gly	Lys 100	Thr	Met	Asp	Tyr	Arg 105	Gly	Thr	Met	Ile	Pro 110	Ser	Glu
25	Ala	Pro	Leu 115	Leu	His	Arg	Gln	Val 120	Lys	Leu	Val	Asp	Pro 125	Met	Asp	Arg
	Lys	Pro 130	Thr	Glu	Ile	Glu	Trp 135	Arg	Phe	Thr	Glu	Ala 140	Gly	Glu	Arg	Va]
30	Arg 145	Val	Ser	Thr	Arg	Ser 150	Gly	Arg	Ile	Ile	Pro 155	Lys	Pro	Glu	Phe	Pro 160
35	Arg	Ala	Asp	Gly	Ile 165	Val	Pro	Glu	Thr	Trp 170	Ile	Asp	Gly	Pro	Lys 175	Asp
	Thr	Ser	Val	Glu 180	Asp	Ala	Leu	Glu	Arg 185	Thr	Tyr	Val	Pro	Cys 190	Leu	Lys
40	Thr	Leu	Gln 195	Glu	Glu	Val	Met	Glu 200	Ala	Met	Gly	Ile	Lys 205	Glu	Thr	Arg
	Lys	Tyr 210	Lys	Lys	Val	Tyr	Trp 215	Tyr								
45	(2)	TNFO	ORMAT	PTON	FOR	SEO	TD N	J∩• 1	35.							
	(2)		(i) :													
50			(xi)	()	A) L B) T D) T	ENGT YPE : OPOL	H: 4 ami: OGY:	9 am no a lin	ino a cid ear	acid		: 13!	5:			
55	Met 1	Ser	Leu	Arg	Gln 5	Lys	Ser	Ser	Phe	Arg 10	Leu	Met	Val	Met	Ser 15	Leu
60	Thr	Ile	Leu	Lys 20	Leu	Ser	Lys	Thr	Thr 25	Val	Leu	Cys	Leu	Arg 30	Cys	Leu

```
His Ser Leu Lys Leu Thr Trp Arg Asp Gly Ala Arg Cys Ile Asn Ala
                                  40
     Glu
 5
      (2) INFORMATION FOR SEQ ID NO: 136:
10
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 68 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
15
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 136:
     Met Ser Gly Ser Phe Ile Leu Cys Leu Ala Leu Val Thr Arg Trp Ser
             5 .
       1
      Pro Gln Ala Ser Ser Val Pro Leu Ala Val Tyr Glu Ser Lys Thr Arg
20
      Lys Ser Tyr Arg Ser Gln Arg Asp Arg Asp Gly Lys Asp Arg Ser Gln
                                 40
              35
25
      Gly Met Gly Leu Ser Leu Leu Val Glu Thr Arg Lys Leu Leu Leu Ser
                             55
      Ala Asn Gln Gly
30
      65
      (2) INFORMATION FOR SEQ ID NO: 137:
35
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 52 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
40
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 137:
      Met Cys Phe Arg Phe Phe Leu Phe Cys Ser Arg Ile Leu Leu Lys Leu
                               10
      Phe Phe Leu Leu Phe Pro Ala Ser Ala Phe Pro Leu Ser Thr Arg Ser
45
      Ser Leu Ser Val Asn Glu His Val Val Val Ser Pro Arg Ser Thr Val
                                   40
                                                       45
 50
      Ser Ile Ser Arg
           50
 55
       (2) INFORMATION FOR SEQ ID NO: 138:
              (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 541 amino acids
 60
                     (B) TYPE: amino acid
```

(D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 138:

5	Met	Val	. Arg	J Thr	Asp 5		His	Thr	Leu	Ser 10		Lys	Arg	ı Asn	Туг 15	Gln
	Val	Thr	Asr	n Ser 20	Met	Phe	Gly	Ala	Ser 25		Lys	Lys	Phe	Val		Gly
10	Val	. Asp	Ser 35	Asp	Tyr	His	Asp	Glu 40		Met	Tyr	Tyr	Ser 45		Ser	Ser
15	Met	Phe 50	Pro	His	Arg	Ser	Glu 55		Asp	Met	Leu	Ala 60	Ser	Pro	Ser	Thr
	Ser 65	Gly	Gln	Leu	Ser	Gln 70	Phe	Gly	Ala	Ser	Leu 75	Tyr	Gly	Gln	Gln	Ser 80
20	Ala	Leu	Gly	' Leu	Pro 85	Met	Arg	Gly	Met	Ser 90	Asn	Asn	Thr	Pro	Gln 95	Leu
	Asn	Arg	Ser	Leu 100	Ser	Gln	Gly	Thr	Gln 105	Leu	Pro	Ser	His	Val 110	Thr	Pro
25	Thr	Thr	Gly 115	Val	Pro	Thr	Met	Ser 120	Leu	His	Thr	Pro	Pro 125	Ser	Pro	Ser
30	Arg	Gly 130	Ile	Leu	Pro	Met	Asn 135	Pro	Xaa	Asn	Met	Met 140	Asn	His	Ser	Gln
	145			Gly		150					155					160
35				Gly	165					170					175	
40				Gln 180					185					190		
40			195	Met				200					205			
45		210		Ile			215					220				
	Leu 225	Asp	Leu	Ser	Asp	Phe 230	Pro	Ala	Leu	Ala	Asp 235	Arg	Asn	Arg	Arg	Glu 240
50	Gly	Ser	Gly	Asn	Pro 245	Thr	Pro	Leu	Ile	Asn 250	Pro	Leu	Ala	Gly	Arg 255	Ala
	Pro	Tyr	Val	Gly 260	Met	Val	Thr	Lys	Pro 265	Ala	Asn	Glu	Gln	Ser 270	Gln	Asp
55	Phe	Ser	Ile 275	His	Asn	Glu	Asp	Phe 280	Pro	Ala	Leu		Gly 285	Ser	Ser	Tyr
60	Lys	Asp 290	Pro	Thr	Ser	Ser	Asn 295	Asp	Asp	Ser		Ser 300	Asn	Leu	Asn	Thr

	Ser 305	Gly	Lys	Thr	Thr	Ser 310	Ser	Thr	Asp	Gly	Pro 315	Lys	Phe	Pro	Gly	Asp 320
5	Lys	Ser	Ser	Thr	Thr 325	Gln	Asn	Asn	Asn	Gln 330	Gln	Lys	Lys	Gly	Ile 335	Gln
	Val	Leu	Pro	Asp 340	Gly	Arg	Val	Thr	Asn 345	Ile	Pro	Gln	Gly	Met 350	Val	Thr
10	Asp	Gln	Phe 355	Gly	Met	Ile	Gly	Leu 360	Leu	Thr	Phe	Ile	Arg 365	Ala	Ala	Glu
15	Thr	Asp 370	Pro	Gly	Met	Val	His 375	Leu	Ala	Leu	Gly	Ser 380	Asp	Leu	Thr	Thr
13	Leu 385	Gly	Leu	Asn	Leu	Asn 390	Ser	Pro	Glu	Asn	Leu 395	Tyr	Pro	Lys	Phe	Ala 400
20	Ser	Pro	Trp	Ala	Ser 405	Ser	Pro	Cys	Arg	Pro 410	Gln	Asp	Ile	Asp	Phe 415	His
	Val	Pro	Ser	Glu 420	Tyr	Leu	Thr	Asn	Ile 425	His	Ile	Arg	Asp	Lys 430	Leu	Ala
25	Ala	Ile	Lys 435	Leu	Gly	Arg	Tyr	Gly 440	Glu	Asp	Leu	Leu	Phe 445		Leu	Tyr
30	Tyr	Met 450		Gly	Gly	Asp	Val 455		Gln	Leu	Leu	Ala 460	Ala	Val	Glu	Leu
30	Phe 465	Asn	Arg	Asp	Trp	Arg 470		His	Lys	Glu	Glu 475	Arg	Val	Trp	Ile	Thr 480
35	Arg	Ala	. Pro	Gly	Met 485	Glu	Pro	Thr	Met	Lys 490		Asn	Thr	Tyr	Glu 495	Arg
	Gly	Thr	Tyr	Туr 500		Phe	Asp	Cys	Leu 505		Trp	Arg	Lys	510		Lys
40	Glu	Phe	His 515		Glu	Tyr	Asp	520		ı Glu	ı Glu	ı Arg	Pro 525		: Leu	Pro
45	Ser	Thr 530		e Asn	Tyr	Asn	535	Ala	Glr	n Glr	n Ala	Phe 540		a		
	(2)	INE	FORMA	MOITA	1 FOF	R SEÇ) ID	NO:	139	:						
50			(i)	SEQ	(A) :	LENG Type	TH: : am	TERIS 58 au ino : li	mino acid	aci	ds					
55			(xi) SE				IPTI			ID N	0: 1	39:			
		z Ile L	e Cys	s Pro		n Cys	s Pro	o Lei	ı Se	r Le		u Cys	s Le	u Ile	e Sei	r Ser 5
60	Le	ı Cy	s Se:	r Le	_	l Ile	e Gl	n Ile	e Se 2		u Ly:	s Th	r Il	e Ar		o Ile

	Thr Leu Leu Asn Met Val Gly Ile Lys Phe Ser Ile Ser Leu Ser A	ısn
5	Lys Ile Asn Ile Asn Ser Arg Thr Trp Xaa 50 55	
10	(2) INFORMATION FOR SEQ ID NO: 140:	
15	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 202 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 140: 	
20	Met Thr Leu Arg Pro Ser Leu Leu Pro Leu His Leu Leu Leu L 1 5 10 15	
	Leu Leu Ser Ala Ala Val Cys Arg Ala Glu Ala Gly Leu Glu Thr G 20 25 30	lu
25	Ser Pro Val Arg Thr Leu Gln Val Glu Thr Leu Val Glu Pro Pro G 35 40 45	lu
	Pro Cys Ala Glu Pro Ala Ala Phe Gly Asp Thr Leu His Ile His Ty 50 55 60	yr
30	Thr Gly Ser Leu Val Asp Gly Arg Ile Ile Asp Thr Ser Leu Thr As 65 70 75	rg 30
35	Asp Pro Leu Val Ile Glu Leu Gly Gln Lys Gln Val Ile Pro Gly Le 85 90 95	eu
	Glu Gln Ser Leu Leu Asp Met Cys Val Gly Glu Lys Arg Arg Ala II 100 105 110	le
40	Ile Pro Ser His Leu Ala Tyr Gly Lys Arg Gly Phe Pro Pro Ser Va 115 120 125	ıl
	Pro Ala Asp Ala Val Val Gln Tyr Asp Val Glu Leu Ile Ala Leu Il 130 135 140	.e
45	Arg Ala Asn Tyr Trp Leu Lys Leu Val Lys Gly Ile Leu Pro Leu Val 145 55 160	
50	Gly Met Ala Met Val Pro Ala Leu Leu Gly Leu Ile Gly Tyr His Le 165 170 175	·u
	Tyr Arg Lys Ala Asn Arg Pro Lys Val Ser Lys Lys Lys Leu Lys Gl 180 185 190	u
55	Glu Lys Arg Asn Lys Ser Lys Lys Lys Xaa 195 200	
60	(2) INFORMATION FOR SEQ ID NO: 141:	

	(1) SEQUENCE CHARACTERISTICS: (A) LENGTH: 217 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear 5 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 141:															
5			(xi)							EQ II	ONO:	141	L:			
	Met 1	Phe	Leu	Arg	Leu 5	Tyr	Leu	Ile	Ala	Arg 10	Val	Met	Leu	Leu	His 15	Ser
10	Lys	Leu	Phe	Thr 20	Asp	Ala	Ser	Ser	Arg 25	Ser	Ile	Gly	Ala	Leu 30	Asn	Lys
15	Ile	Asn	Phe 35	Asn	Thr	Arg	Phe	Val 40	Met	Lys	Thr	Leu	Met 45	Thr	Ile	Cys
13	Pro	Gly 50	Thr	Val	Leu	Leu	Val 55	Phe	Ser	Ile	Ser	Leu 60	Trp	Ile	Ile	Ala
20	Ala 65	Trp	Thr	Val	Arg	Val 70	Cys	Glu	Ser	Pro	Glu 75	Ser	Pro	Ala	Gln	Pro 80
	Ser	Gly	Ser	Ser	Leu 85	Pro	Ala	Trp	Tyr	His 90	Asp	Gln	Gln	Asp	Va1 95	Thr
25	Ser	Asn	Phe	Leu 100	Gly	Ala	Met	Trp	Leu 105	Ile	Ser	Ile	Thr	Phe 110	Leu	Ser
20	Ile	Gly	Tyr 115	Gly	Asp	Met	Val	Pro 120	His	Thr	Tyr	Cys	Gly 125	Lys	Gly	Val
30	Cys	Leu 130		Thr	Gly	Ile	Met 135		Ala	Gly	Суѕ	Thr 140		Leu	Val	Val
35	Ala 145		. Val	Ala	Arg	Lys 150		Glu	Leu	Thr	Lys 155		Glu	Lys	His	Val 160
	His	Asr	n Phe	Met	Met 165		Thr	Gln	Leu	Thr 170		Arg	Ile	Lys	Asn 175	
40	Ala	. Ala	a Asn	Val 180		. Arg	g Glu	Thr	Trp		ı Ile	тут	Lys	His 190		Lys
45	Leu	Le.	1 Lys 195		: Ile	e Asp	His	Ala 200		: Val	L Arg	Lys	His 205	Glr	Arg	, Lys
45	Ph∈	210		Ser	туг	r Pro	215		. Xaa	ı						
50	(2)	IN	FORM/	10ITA	1 FOE	R SE(Q ID	NO:	142	:						
			(i)	SEQ			ARAC'				ids					
55			(xi) SE	(D)	TOPO	: am LOGY ESCR	: li	near		ID N	0: 1	42:			
60		t Se 1	r As	n Th		r Va 5	l Pr	o As	n Al		o Gl 0	n Al	a As	n Se	r As	

	Met	: Val	. Gly	туr 20	Val	. Leu	Gly	Pro	Phe 25		Leu	ıle	Thr	Leu 30		. Gly
5	Va]	. Val	. Val 35	Ala	Val	Val	Met	Туг 40	Val	Gln	Lys	Lys	Lys 45		Val	Asp
10	Arg	Leu 50	Arg	His	His	Leu	Leu 55	Pro	Met	Tyr	Ser	Туг 60		Pro	Ala	Glu
	Glu 65	Leu	His	Glu	Ala	Glu 70	Gln	Glu	Leu	Leu	Ser 75		Met	Gly	Asp	Pro 80
15	Lys	Val	Val	His	Gly 85	Trp	Gln	Ser	Gly	Tyr 90	Gln	His	Lys	Arg	Met 95	Pro
	Leu	Leu	Asp	Val 100	Lys	Thr										
20																
	(2)	INF		TION												
25				(A) L B) T D) T	ENGT YPE: OPOL	H: 1 ami OGY:	12 a no a lin	mino cid ear	aci		. 14	٦.			
30	35															
50	мес 1	Arg	Giu	Cys	Gin 5	GIU	GIu	Ser	Phe	Trp 10	Lys	Arg	Ala	Leu	Pro 15	Phe
35	Ser	Leu	Val	Ser 20	Met	Leu	Val	Thr	Gln 25	Gly	Leu	Val	Tyr	Gln 30	Gly	Tyr
	Leu	Ala	Ala 35	Asn	Ser	Arg	Phe	Gly 40	Ser	Leu	Pro	Lys	Val 45	Ala	Leu	Ala
40	Gly	Leu 50	Leu	Gly	Phe	Gly	Leu 55	Gly	Lys	Val	Ser	Tyr 60	Ile	Gly	Val	Cys
	Gln 65	Ser	Lys	Phe	His	Phe 70	Phe	Glu	Asp	Gln	Leu 75	Arg	Gly	Ala	Gly	Phe 80
45	Gly	Pro	Gln	His	Asn 85	Arg	His	Cys	Leu	Leu 90	Thr	Cys	Glu	Glu	Cys 95	Lys
50	Ile	Lys	His	Gly 100	Leu	Ser	Glu	Lys	Gly 105	Asp	Ser	Gln	Pro	Ser 110	Ala	Ser
55	(2)	INFC	RMAT	'ION	FOR	SEQ	ID N	lO: 1	44:							
		((i) S	EQUE												
60						INGTH				cids	ł					

	(D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 144: Met Lys Asn Asp Arg Asn Gln Gly Phe Ser Leu Leu Gln Leu Ile Asp 1 5 10 15 Trp Asn Lys Pro															
5		Lys .	Asn ,	Asp A		Asn (Gln (Gly :	Phe S		Leu 1	Leu ·	Gln	Leu		Asp
	Trp	Asn i	Lys	Pro 20												
10																
	(2)	INFO	RMAT	ION I	FOR	SEQ	ID N	0: 1	45:							
15				(E	L) LE 3) TY 0) TO	INGTH (PE: OPOLO	H: 30 amir XGY:) ami no ac line	no a rid ear			145	5:			
20	Met 1	Gly	Thr	Gln	Pro 5	Pro	Val	Val	Ala	Gly 10	Phe	Thr	Ile	Pro	Met 15	Leu
25	Gly	Tyr	Thr	Val 20	Arg	Val	Leu	Thr	Phe 25	His	Leu	Ser	Cys	Ser 30		
	(2)	INFO	RMAT	NOI	FOR	SEQ	ID 1	10: 1	.46:							
30			(i) :	(1		ENGT: YPE :	H: 9 ami	9 am no a	ino a		s					
35				SEQ												
	Met 1	Lys	Ile	Pro	Val 5	Leu	Pro	Ala	Val	Va1 10	Leu	Leu	Ser	Leu	Leu 15	Val
40	Leu	His	Ser	Ala 20	Gln	Gly	Ala	Thr	Leu 25	Gly	Gly	Pro	Glu	G1u 30	Glu	Ser
	Thr	Ile	Glu 35	Asn	Tyr	Ala	Ser	Arg 40	Pro	Glu	Ala	Phe	Asn 45	Thr	Pro	Phe
45	Leu	Asn 50		Asp	Lys	Leu	Arg 55	Ser	Ala	Phe	Lys	Ala 60		Glu	Phe	Leu
50	Asn 65	_	His	Ala	Leu	Phe 70		Ser	Ile	Lys	Arg 75		: Leu	ı Pro	Phe	Let 80
	Asn	Trp	Asp	Ala	Phe 85		Lys	Leu	Lys	Gly 90		Arg	sei	c Ala	Thr 95	
55	Asp	Ala	Gln	1												
60	(2)	INE	FORM	MOIT!	FOF	SEÇ	Q ID	NO:	147:							

```
(i) SEQUENCE CHARACTERISTICS:
                      (A) LENGTH: 8 amino acids
                      (B) TYPE: amino acid
                      (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 147:
       Met Val Trp Gly Leu Leu Gly
                        5
 10
       (2) INFORMATION FOR SEQ ID NO: 148:
              (i) SEQUENCE CHARACTERISTICS:
 15
                     (A) LENGTH: 39 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 148:
20
      Met Leu Pro Leu Leu Ser Leu Leu Phe Leu Phe Phe Ser Thr Val Ser
                                          10
      Ser Phe Cys Gly Met Pro Leu Arg Ala His Thr Arg Ala Xaa Ala His
                   20
                                       25
25
      Thr Arg Thr Phe Ala Ser Arg
               35
30
       (2) INFORMATION FOR SEQ ID NO: 149:
              (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 131 amino acids
35
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 149:
      Met Ile Cys Glu Thr Lys Ala Arg Lys Ser Ser Gly Gln Pro Gly Arg
40
      Leu Pro Pro Pro Thr Leu Ala Pro Pro Gln Pro Pro Leu Pro Glu Thr
                                       25
45
      Ile Glu Arg Pro Val Gly Thr Gly Ala Met Val Ala Arg Ser Ser Asp
      Leu Pro Tyr Leu Ile Val Gly Val Val Leu Gly Ser Ile Val Leu Ile
                               55
                                                   60
50
      Ile Val Thr Phe Ile Pro Phe Cys Leu Trp Arg Ala Trp Ser Lys Gln
      Lys His Thr Thr Asp Leu Gly Phe Pro Arg Ser Ala Leu Pro Pro Ser
55
                                         90
      Cys Pro Tyr Thr Met Val Pro Leu Gly Gly Leu Pro Gly His Gln Ala
                                      105
60
      Val Asp Ser Pro Thr Ser Val Ala Ser Val Asp Gly Pro Val Leu Met
```

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120 125 115 Gly Ser Thr 130 5 (2) INFORMATION FOR SEQ ID NO: 150: 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 32 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 150: 15 Met Gly Ala Pro Ser Leu Thr Met Leu Leu Leu Leu Lys Val Gln Pro Arg Arg Thr Gln Ala Phe Asp Ala His Trp Val Gly Leu Pro Leu Leu 20 25 25 (2) INFORMATION FOR SEQ ID NO: 151: (i) SEQUENCE CHARACTERISTICS: 30 (A) LENGTH: 14 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 151: Met Cys Leu Ile Phe Leu Leu Leu Leu Leu Ser Phe Ser 35 5 10 1 40 (2) INFORMATION FOR SEQ ID NO: 152: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 8 amino acids (B) TYPE: amino acid 45 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 152: His Pro His Gln Asp Ser Gln Pro 1 5 50 (2) INFORMATION FOR SEQ ID NO: 153: (i) SEQUENCE CHARACTERISTICS: 55 (A) LENGTH: 68 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 153: 60

	1			Ser	5		. nea	ALG	Leu	10		vai	. vai	. Ser	15	
5	Ile	Tyr	Leu	Ala 20		His	Pro	Leu	Leu 25	Ser	Phe	Ser	Leu	Glu 30	Ser	Pro
	Leu	Leu	Val 35		Trp	Arg	Asp	Cys 40		Gln	Asn	Ile	Trp 45		Ser	Gly
10	Ser	Val 50	Trp	Tyr	Lys	Arg	Trp 55		Leu	Pro	His	Met 60		Val	Cys	Cys
15	Gln 65	Asp	Leu	His												
•	(2)				FOR											
20			(i)	(ENCE A) L B) T	ENGT	Ή: 2	6 am	ino		ls					
25			(xi)		D) T UENC					EQ I	D NO	: 15	4:			
	Met 1	Leu	Lys	Ile	Phe 5	Lys	Glu	Trp	Glu	Asn 10	Leu	Asn	Leu	Ile	Leu 15	Thr
30	Ser	Ile	Arg	Ile 20	Leu	Glu	Arg	Gln	Asn 25	Met						
35	(2)	INFO	ORMA!	rion	FOR	SEQ	ID 1	NO: 1	155:							
40			(i) :	(. (.	ENCE A) L B) T D) T	ENGT YPE :	H: 1 ami	95 a no a	mino cid		ds					
40					JENCI											
	Met 1	Asp	Cys	Glu	Val 5	Asn	Asn	Gly	Ser	Ser 10	Leu	Arg	Asp	Glu	Cys 15	Ile
45	Thr	Asn	Leu	Leu 20	Val	Phe	Gly	Phe	Leu 25	Gln	Ser	Cys	Ser	Asp 30	Asn	Ser
50	Phe	Arg	Arg 35	Glu	Leu	Asp	Ala	Leu 40	Gly	His	Glu	Leu	Pro 45	Val	Leu	Ala
	Pro	Gln 50	Trp	Glu	Gly	Tyr	Asp 55	Glu	Leu	Gln	Thr	Asp 60	Gly	Asn	Arg	Ser
55	Ser 65	His	Ser	Arg	Leu	Gly 70	Arg	Ile	Glu	Ala	Asp 75	Ser	Glu	Ser	Gln	Glu 80
	Asp	Ile	Ile	Arg	Asn 85	Ile	Ala	Arg	His	Leu 90	Ala	Gln	Val	Gly	Asp 95	Ser
50	Met	Asp	Arg	Ser	Ile	Pro	Pro	Gly	Leu	Val	Asn	Gly	Leu	Ala	Leu	Gln

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				100					105					110		
_	Leu	Arg	Asn 115	Thr	Ser	Arg	Ser	Glu 120	Glu	Asp	Arg	Asn	Arg 125	Asp	Leu	Ala
5	Thr	Ala 130	Leu	Glu	Gln	Leu	Leu 135	Gln	Ala	Tyr	Pro	Arg 140	Asp	Met	Glu	Lys
10	Glu 145	Lys	Thr	Met	Leu	Val 150	Leu	Ala	Leu	Leu	Leu 155	Ala	Lys	Lys	Val	Ala 160
	Ser	His	Thr	Pro	Ser 165	Leu	Leu	Arg	Asp	Val 170	Phe	His	Thr	Thr	Val 175	Asn
15	Phe	Ile	Asn	Gln 180	Asn	Leu	Arg	Thr	Tyr 185	Val	Arg	Ser	Leu	Ala 190	Arg	Asn
20	Gly	Met	Asp 195													
	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	156:							
25			(i)		(A) I (B) :	LENG!	TH: !		mino acid		ls					
30			•	SEÇ	-									mi		
	Met 1		. Leu	ı Ser	Leu		. Sei	. Val	l Ser	7 Val		r Pro	Ser	r Thr	15	Ala
35	Cys	: Ser	: Phe	Let 20		Pro	Lys	s Ala	a Arg		Ser	: Lys	s Arg	Ser 30		Arg
	Asr	туі	c Thi		Sei	Thi	s Sei	r Pro		y Gly	y Pro	Arq	g Ála 49		Arg	g Gly
40	Gly	Ala 50		o Arg	g Lei	ı Sei	r Se: 5!		n Gli	n Ası	n Sei	r Sei 60		o Lys	s Gly	y Val
45	Ala 6		l Ala	a Ly:	s Ala	a Se:		r Ar	g Pr	o Va	1 Le: 7:	_	s Ph	e Lei	ı Pro	Gly 80
	Pr	o Tr	p Se	r Se	r Xa 8	_	o Xa	a Al	a Ph	e Le		e				
50	(2) IN	FORM	ATIO	N FO	R SE	Q IE	NO:	157	:						
55				SEÇ L) SI	(A) (B) (D)	LENO TYPE TOPO	ETH: E: an OLOG	31 a mino Y: 1:	amino acio inea	o aci d r		10: 1	L57 :			
60	Me	et Gl	Ly Th	ır Le	eu Se	er Al	.a Gl	lu Cy	s Se		y Pr LO	o Al	a Th	ır Le		y Leu .5

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Cys Leu Val Val Pro Trp Asn Ser Ser Gly Leu Ser Gln Pro Pro
                   20
                                       25
 5
      (2) INFORMATION FOR SEQ ID NO: 158:
             (i) SEQUENCE CHARACTERISTICS:
10
                     (A) LENGTH: 91 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 158:
15
      Met Lys Phe Leu Ala Val Leu Val Leu Gly Val Ser Ile Phe Leu
        1
      Val Ser Ala Gln Asn Pro Thr Thr Ala Ala Pro Ala Asp Thr Tyr Pro
                                       25
20
      Ala Thr Gly Pro Ala Asp Asp Glu Ala Pro Asp Ala Glu Thr Thr Ala
      Ala Ala Thr Thr Ala Thr Thr Ala Ala Pro Thr Thr Ala Thr Thr Ala
25
      Ala Ser Thr Thr Ala Arg Lys Asp Ile Pro Val Leu Pro Lys Trp Val
                          70
30
      Gly Asp Leu Pro Asn Gly Arg Val Cys Pro Xaa
35
      (2) INFORMATION FOR SEQ ID NO: 159:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 89 amino acids
                    (B) TYPE: amino acid
40
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 159:
      Met Ile Ile Ser Leu Phe Ile Tyr Ile Phe Leu Thr Cys Ser Asn Thr
45
      Ser Pro Ser Tyr Gln Gly Thr Gln Leu Gly Leu Gly Leu Pro Ser Ala
                   20
                                       25
      Gln Trp Trp Pro Leu Thr Gly Arg Arg Met Gln Cys Cys Arg Leu Phe
50
      Cys Phe Leu Leu Gln Asn Cys Leu Phe Pro Phe Pro Leu His Leu Ile
           50
55
      Gln His Asp Pro Cys Glu Leu Val Leu Thr Ile Ser Trp Asp Trp Ala
                          70
      Glu Ala Gly Ala Ser Leu Tyr Ser Pro
                      85
60
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	(2)	INFO	RMAT	ION	FOR	SEQ	ID N	0: 1	60:							
5			(i) S	(F (E	A) L1 3) T' 5) T(ENGTI YPE : OPOLO	H: 17 amir DGY:	74 am no ac line	nino :id ear	ació						
10	Met 1		(xi) Ser											Val	Leu 15	Ser
15	Phe	Val	Phe	Gly 20	Cys	Asn	Val	Leu	Arg 25	Ile	Leu	Leu	Pro	Ser 30	Phe	Ser
	Ser	Phe	Met 35	Ser	Arg	Val	Leu	Gln 40	Lys	Asp	Ala	Glu	Gln 45	Glu	Ser	Gln
20	Met	Arg 50	Ala	Glu	Ile	Gln	Asp 55	Met	Lys	Gln	Glu	Leu 60	Ser	Thr	Val	Asn
25	65		Asp			70					75					80
	_		Thr		85					90					95	
30				100					105					110		Gln
			115					120					125			Val
35		130	1				135					140				Ala
40	145					150	1				155					Leu 160
	Val	. Cys	s Asn	. Lys	Val 165		. Ala	ı Ile	Val	. Leu 170		: Pro) Phe	Ser		
45	(2)	INE	FORMA	ATION	1 FOI	R SEÇ	Q ID	NO:	161:	1						
50			(i)		(A)	LENG	TH:	TERIS 45 au ino a	mino	aci	ds					
50			(xi) SE	(D)	торо	LOGY	: li	near		ID N	0: 1	61:			
55		t Gl; 1	y Ly:	s Le		e Asi 5	n Il	e Val	l Il	e Arg		s Pro	o Lei	ı Le	u Lei 1	ı Leu 5
	Le:	u Va	l Gli	n Cys		u As	n Cy	s Cy:	s Ar		s As:	n Me	t Le	и Ту: 3		n Ile
60	Ph	e Le	u As	n Il	e Hi	s As	n Il	e Hi	s Ly	s Ph	e Se	r As	n Hi	s		

35 40 45 5 (2) INFORMATION FOR SEQ ID NO: 162: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 amino acids (B) TYPE: amino acid 10 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 162: Met Val Ala Ser Thr Leu Val Thr Asn Leu Phe Gly Val Ala Phe Ala 10 15 Thr Thr Ala Ala Thr Arg Ala 20 20 (2) INFORMATION FOR SEQ ID NO: 163: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 70 amino acids 25 (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 163: Met Leu Met Ala Pro Val Val Cys Leu Ser Phe Ser Pro Cys Pro Ala 30 Asp Thr Ser Leu Thr Gly Asp Gly Leu Lys Ala Gly Leu Glu Arg Gly 35 Xaa Ala Leu Val Thr Leu Phe Asp Ser Val Thr His Phe Leu Ala His Thr Leu Phe Glu Leu Leu Asp Phe Gln Leu Ala Phe Leu Arg Ser Gly 55 40 Lys Gln Thr Ala Pro His 65 45 (2) INFORMATION FOR SEQ ID NO: 164: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 323 amino acids 50 (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 164: Met Leu Leu Leu Leu Leu Gly Ser Gly Gln Gly Pro Gln Gln 55 1 Val Gly Ala Gly Gln Thr Phe Glu Tyr Leu Lys Arg Glu His Ser Leu

Ser Lys Pro Tyr Gln Gly Val Gly Thr Gly Ser Ser Ser Leu Trp Asn

			35					40					45			
<i>-</i>	Leu	Met 50	Gly	Asn	Ala	Met	Val 55	Met	Thr	Gln	Tyr	Ile 60	Arg	Leu	Thr	Pro
5	Asp 65	Met	Gln	Ser	Lys	Gln 70	Gly	Ala	Leu	Trp	Asn 75	Arg	Val	Pro	Cys	Phe 80
10	Leu	Arg	Asp	Trp	Glu 85	Leu	Gln	Val	His	Phe 90	Lys	Ile	His	Gly	Gln 95	Gly
	Lys	Lys	Asn	Leu 100	His	Gly	Asp	Gly	Leu 105	Ala	Ile	Trp	Tyr	Thr 110	Arg	Asn
15	Arg	Met	Gln 115	Pro	Gly	Pro	Val	Phe 120	Gly	Asn	Met	Asp	Lys 125	Phe	Val	Gly
20	Leu	Gly 130	Val	Phe	Val	Asp	Thr 135	Tyr	Pro	Asn	Glu	Glu 140	Lys	Gln	Gln	Glu
20	Arg 145	Val	Phe	Pro	Tyr	Ile 150	Ser	Ala	Met	Val	Asn 155	Asn	Gly	Ser	Leu	Ser 160
25					165					170				Gly	175	
	Ala	Ile	· Val	Arg 180		Leu	His	Tyr	Asp 185		Phe	Leu	Val	11e 190	Arg	Tyr
30		_	195	•				200					205			
35		210)				215	•				220		Arg		
	225	5				230)				235					Asp 240
40					245	ò				250)				255	
				260)				265	5				270)	Met
45			27	5				280)				285	5		ı Ala
50		29	0				29	5				300)			a Ile
	Va 30		e Gl	y Il	e Il	e Le		r Ası	n Ly:	s Tr	p Gl: 31		u Gl	n Sei	r Arg	320
55	Ar	g Ph	е Ту	r												
<i>(</i> 0	(2	()	FORM	iatic	N FO	R SE	Q II	NO:	165	:						
60																

			(1)	SEQ		LENG	TH:	321 ino	amin		ids					
5			(xi		(D)	TOPO:	LOGY	: lii IPTIO	near	SEQ :	ID NO	D: 16	55:			
	Met	Pro	Se ₁	Glu	ı Туг 5	Thr	туз	. Val	. Lys	Let		g Ser	Asr	Cys	Ser 15	Arg
10	Pro	Ser	Lev	Glr 20	Trp	Tyr	Thr	Arg	Ala 25		Ser	Lys	Met	Arg		Pro
15	Ser	Leu	Leu 35	Leu	Lys	Asp	· Ile	Leu 40	Lys	Cys	Thr	· Leu	Leu 45		Phe	Gly
	Val	Trp 50) Il∈	: Leu	Tyr	Ile	Leu 55	Lys	Leu	Asn	Tyr	Thr 60		Glu	Glu	Cys
20	Asp 65	Met	Lys	Lys	Met	His 70	Tyr	Val	Asp	Pro	Asp 75		Val	Lys	Arg	Ala 80
	Gln	Lys	Tyr	Ala	Gln 85	Gln	Val	Leu	Gln	Lys 90	Glu	Cys	Arg	Pro	Lys 95	Phe
25	Ala	Lys	Thr	Ser 100	Met	Ala	Leu	Leu	Phe 105	Glu	His	Arg	Tyr	Ser 110	Val	Asp
30	Leu	Leu	Pro 115	Phe	Val	Gln	Lys	Xaa 120	Pro	Lys	Asp	Ser	Glu 125	Ala	Glu	Ser
	Lys	Tyr 130	Asp	Pro	Pro	Phe	Gly 135	Phe	Arg	Lys	Phe	Ser 140	Ser	Lys	Val	Gln
35	Thr 145	Leu	Leu	Glu	Leu	Leu 150	Pro	Glu	His	qzA	Leu 155	Pro	Glu	His	Leu	Lys 160
	Ala	Lys	Thr	Cys	Arg 165	Arg	Суѕ	Val	Val	Ile 170	Gly	Ser	Gly	Gly	Ile 175	Leu
40	His	Gly	Leu	Glu 180	Leu	Gly	His	Thr	Leu 185	Asn	Gln	Phe	Asp	Val 190	Val	Ile
45	Arg	Leu	Asn 195	Ser	Ala	Pro	Val	Glu 200	Gly	Tyr	Ser	Glu	His 205	Val	Gly	Asn
	Lys	Thr 210	Thr	Ile	Arg	Met	Thr 215	Tyr	Pro	Glu	Gly	Ala 220	Pro	Leu	Ser	Asp
50	Leu 225	Glu	Tyr	Tyr	Ser	Asn 230	Asp	Leu	Phe	Val	Ala 235	Val	Leu	Phe	Lys	Ser 240
	Val	Asp	Phe	Asn	Trp 245	Leu	Gln	Ala	Met	Val 250	Lys	Lys	Glu	Thr	Leu 255	Pro
55	Phe	Trp	Val	Arg 260	Leu	Phe	Phe	Trp	Lys 265	Gln	Val	Ala	Glu	Lys 270	Ile	Pro
50	Leu	Gln	Pro 275	Lys	His	Phe	Arg	Ile 280	Leu	Asn	Pro	Val	Ile 285	Ile	Lys	Glu

	Thr	Ala 290	Phe	Xaa	His	Pro	Ser 295	Val	Leu	Arg	Ala	Ser 300	Val	Lys	Val	Leu
5	Gly 305	Ala	Glu	Ile	Arg	Thr 310	Ser	Pro	Gln	Ser	Val 315	Ser	Leu	Pro	Leu	Ser 320
	Xaa															
10																
	(2)	INF	ORMA	TION	FOR	SEQ	ID I	: 01/	166:							
15				(A) L B) T D) T	ENGT YPE : OPOL	H: 3 ami OGY:	l am no a lin	nino ncid near	acid		: 16	6:			
20	Met 1		Leu	. Asp	Val	Gln	Thr	Val	Val	Val 10		Ala	Val	Ile	Val 15	Val
25	Leu	ı Lev	ı Lev	val 20		Val	Ile	Leu	Met 25	Phe	Phe	Leu	Gly	Thr 30	Arg	
	(2)	INE	FORMA	MOITA	I FOR	SEQ	ID	NO:	167:							
30			(i)		(A) I (B) 1	LENG TYPE	rH: ' : am:	72 ar ino a	mino acid		ds					
35			(xi) SE((D) (SEQ :	ID N	D: 1	67:			
		t Le	u Pr	o Lev	ı Lev		e Cys	s Ala	a Phe	Cys		ı His	s Lys	s Lei	15	Pro
40	Le	u Le	u Ph	e Lei 20		Asp	Va:	l Le	u Met 25		a His	s Gl	u Ala	a Val		Arg
	Th	r Hi	s Gl 3		e Glı	n Leu	ı Pro	o Ası		Gl:	u Phe	e Pr	o Se:		n Glr	n Asn
45	Gl		l Le O	u Asi	n Ly:	s Thi	r Le		e Ası	n Ly:	s Le	u Ly 6		s Ly:	s Lys	s Lys
50		s Ly 5	rs Ly	rs Xa	a Xa	a Xaa		s Ly	s							
	(2	(!		(ATIO												
55			(i)) SEÇ	(A) (B)	LENC	TH: E: ar	282 mino	STIC amir acid inear	no ao 1	cids					
60			(x	i) SI							ID 1	10:	168:			

	Met 1		Ser	Arg	Gly 5		Arg	Pro	Glu	His 10	Gly	Gly	Pro	Pro	Glu 15	Leu
5	Ph∈	туг	Asp	Glu 20		Glu	Ala	Arg	Lys 25		Val	Arg	Asn	Ser 30	Arg	Met
	Ile	: Asp	Ile 35	Gln	Thr	Arg	Met	Ala 40	Gly	Arg	Ala	Leu	Glu 45	Leu	Leu	Tyr
10	Leu	Pro 50	Glu	Asn	Lys	Pro	Cys 55	Tyr	Leu	Leu	Asp	Ile 60	Gly	Cys	Gly	Thr
15	Gly 65	Leu	Ser	Gly	Ser	Туr 70	Leu	Ser	Asp	Glu	Gly 75	His	Tyr	Trp	Val	Gly 80
	Leu	Asp	Ile	Ser	Pro 85	Ala	Met	Leu	Asp	Glu 90	Ala	Val	Asp	Arg	Glu 95	Ile
20	Glu	Gly	Asp	Leu 100	Leu	Leu	Gly	Asp	Met 105	Gly	Gln	Gly	Ile	Pro 110	Phe	Lys
	Pro	Gly	Thr 115	Phe	Asp	Gly	Cys	Ile 120	Ser	Ile	Ser	Ala	Val 125	Gln	Trp	Leu
25	Cys	Asn 130	Ala	Asn	Lys	Lys	Ser 135	Glu	Asn	Pro	Ala	Lys 140	Arg	Leu	Tyr	Cys
30	145		Ala			150					155					160
			Leu		165					170					175	
35			Thr	180					185					190		
40			Ala 195					200					205			
40		210	Phe				215					220				
45	225		Glu			230					235					240
			Gly		245					250					255	
50			His	260					265		Arg	Pro	Asp	Thr 270	Gln	Tyr
5 5	Thr	Gly	Arg 275	Lys	Arg	Lys	Pro	Arg 280	Phe	Xaa						
55	(2)	INFO	RMAT	'ION	FOR	SEQ	ID N	ю: 1	69:							
60			(i) S	EQUE	NCE	CHAR	ACTE	ERIST	ICS:	acids						

PCT/US98/05311 WO 98/42738

		(:	xi) :	(D) TY!) TO! ENCE	POLO	GY:	line	ar	Q ID	NO:	169	:			
5	Met L	eu (Gly I	Lys 7	Thr I 5	ys I	Phe (Gln s	Ser S	Tyr I 10	lys :	Ser 1	Phe :	Ser 2	Arg 15	Lys
10	Leu M	1et '	Val (Cys 1 20	Pro S	Ser 1	Thr									
15	(2)	(RMAT: i) S	EQUE A) E) I)	NCE (A) LE B) TY D) TO	CHAR NGTH PE:	ACTE 1: 32 amir XGY:	RIST 28 am no ac line	ICS: mino id ar	ació		170):			
20	Met '													Arg	His 15	Gly
25	Ala	Gln	Gly	Lys 20	Pro	Ser	Pro	Asp	Ala 25	Gly	Pro	His	Gly	Gln 30	Gly	Arg
	Val :	His	Gln 35	Ala	Ala	Pro	Leu	Ser 40	Asp	Ala	Pro	His	Asp 45	Asp	Ala	His
30	Gly	Asn 50	Phe	Gln	Tyr	Asp	His 55	Glu	Ala	Phe	Leu	Gly 60	Arg	Glu	Val	Ala
35	Lys 65	Glu	Phe	Asp	Gln	Leu 70	Thr	Pro	Glu	Glu	Ser 75	Gln	Ala	Arg	Leu	Gly 80
	Arg	Ile	Val	Asp	Arg 85	Met	Asp	Arg	Ala	Gly 90	Asp	Gly	Asp	Gly	Trp 95	Val
40	Ser	Leu	Ala	Glu 100	Leu	Arg	Ala	Trp	Ile 105	Ala	His	Thr	Gln	Gln 110	Arg	His
	Ile	Arg	Asp 115	Ser	Val	Ser	Ala	Ala 120	Trp	Asp	Thr	Tyr	Asp 125	Thr	Asp	Arg
45	Asp	Gly 130	Arg	Val	Gly	Trp	Glu 135		Leu	Arg	Asn	Ala 140		Tyr	Gly	His
50	Tyr 145	Ala	Pro	Gly	Glu	Glu 150	Phe	His	Asp	Val	Glu 155		Ala	. Glu	Thr	Туr 160
	Lys	Lys	Met	Leu	Ala 165	Arg	Asp	Glu	Arg	Arg 170		Arg	Val	Ala	175	
55	Asp	Gly	Asp	Ser 180		Ala	Thr	Arg	Glu 185		Leu	Thr	Ala	Phe 190		ı His
	Pro	Glu	195		Pro	His	Met	200) Ile	val	. I1e	205		Thi	r Leu
60	Glu	Asp	Leu	. Asp	Arg	Asn	Lys	s Asp	Gl3	у Туг	· Val	l Glr	ı Val	l Glu	ı Gl	тут

		210					215					220				
5	Ile 225	Ala	Asp	Leu	Tyr	Ser 230	Ala	Glu	Pro	Gly	Glu 235	Glu	Glu	Pro	Ala	Trp 240
	Val	Gln	Thr	Glu	Arg 245	Gln	Gln	Phe	Arg	Asp 250	Phe	Arg	Asp	Leu	Asn 255	Lys
10	Asp	Gly	His	Leu 260	Asp	Gly	Ser	Glu	Val 265	Gly	His	Trp	Val	Leu 270	Pro	Pro
	Ala	Gln	Asp 275	Gln	Pro	Leu	Val	Glu 280	Ala	Asn	His	Leu	Leu 285	His	Glu	Ser
15	Asp	Thr 290	Asp	Lys	Asp	Gly	Arg 295	Leu	Ser	Lys	Ala	Xaa 300	Ile	Leu	Gly	Asn
20	Trp .	Asn	Met	Phe	Val	Gly 310	Ser	Gln	Ala	Thr	Asn 315	Tyr	Gly	Glu	Asp	Leu 320
	Thr A	Arg	His	His	Asp 325	Glu	Leu	Xaa								
25	(2)	INFC	RMAT	NOI	FOR	SEQ	ID N	Ю: 1	.71:							
30				() () ()	A) L1 B) T C) T(CHAF ENGTI YPE: DPOLO	H: 6 amin XGY:	9 am: no ao line	ino a cid ear	acids		: 171	L:			
35	Met (Cys	Trp	Leu	Arg 5	Ala	Trp	Xaa	Gln	Ile 10	Xaa	Leu	Pro	Va1	Phe 15	Xaa
	Ser }	Kaa	Phe	Leu 20	Ile	Gln	Leu	Leu	Ile 25	Ser	Phe	Ser	Glu	Asn 30	Gly	Phe
40	Ile H	lis	Ser 35	Pro	Arg	Asn	Asn	Gln 40	Lys	Pro	Arg	Asp	Gly 45	Asn	Xaa	Glu
45	Glu C	уs . 50	Ala	Val	Lys	Lys	Ser 55	Cys	Gln	Leu	Cys	Thr 60	Glu	Asp	Lys .	Lys
	Tyr M 65	[et]	Met .	Asn .	Arg											
50	(2) I	NFO	RMAT:	ION :	FOR	SEQ :	ID N	0: 1	72:							
55				(A (B (D	L) LE () TY () TC	CHAR INGTH PE: POLO DES	: 16 amir GY:	0 am 0 ac line	ino id ar			172				
60	Met T													Asp .	Ala 1 15	Met

	Val	Met	Asp	Glu 20	Lys	Val	Lys	Arg	Ser 25	Pne	vai	Leu	Asp	30	Ald	Sel
5	Ala	Ile	Cys 35	Asn	Tyr	Asn	Ala	His 40	Tyr	Lys	Asn	His	Pro 45	Lys	Tyr	Trp
10	Cys	Arg 50	Gly	Tyr	Phe	Arg	Asp 55	Tyr	Cys	Asn	Ile	Ile 60	Ala	Phe	Ser	Pro
10	Asn 65	Ser	Thr	Asn	His	Val 70	Ala	Leu	Lys	Asp	Thr 75	Gly	Asn	Gln	Leu	Ile 80
15	Val	Thr	Met	Ser	Cys 85	Leu	Asn	Lys	Glu	Asp 90	Thr	Gly	Trp	Tyr	Trp 95	Cys
	Gly	Ile	Gln	Arg 100	Asp	Phe	Ala	Arg	Asp 105	Asp	Met	Asp	Phe	Thr 110	Glu	Leu
20	Ile	Val	Thr 115		Asp	Lys	Gly	Thr 120	Trp	Pro	Met	Thr	Leu 125	Val	Trp	Glu
25	Arg	Leu 130		Gly	Thr	Lys	Pro 135	Glu	Ala	Ala	Arg	Leu 140		Lys	Leu	Ser
23	Ala 145		Leu	Thr	Ala	Pro 150	Gly	Arg	Pro	Phe	Ser 155		Phe	Ala	Tyr	Xaa 160
30																
35	(2)	INF		SEQU	JENCE (A) I (B) T	CHA LENGT	RACT TH: :	TERIS 123 a ino a		3:	ids					
40			(xi		(D) I				N: S	SEQ :	ID N	o: 17	73:			
	Met		a Xaa	a His	Phe 5		Le:	ı Val	Ala	Lei		n Ser	Val	. Pro	His 15	Cys
45	Pro) Hi	s Lei	Let 20		Glu	ı Glu	ı His	Lys 25		ı Cy:	s Lys	s Val	Ser 30		Phe
50	Se	r Gl	y Va 3		r Leu	ı Val	L Thi	c Sea		g Gl	n As	p Sei	r Sei 49		туг	· Val
30	Pr	o Va 5		n Thi	r Lei	ı Phe	e Ile 5	_	s Le	u Gl	y Pr	o Trj		a Trị	Asp	Leu
55	Xa 6		о Су	s Thi	r Ala	a Gli 70		p Pr	o Gl	u Al	a Gl 7		g Se	r Lei	u Arg	g Leu 80
	Су	s Hi	s Se	r Hi	s Le		a Ar	g Xa	a As		1 Se 0	r Pr	o Se	r Gl	n Ala	a Ala
60	Gl	u Gl	у Ха	a Xa	a Xa	a Ar	g Gl	у Су	s Gl	n Hi	s Ar	g Gl	y Se	r Ar	g Gl	ı Leu

				100					105					110		
5	Thr	Phe	Leu 115	Ser	Ala	Glu	Asn	Glu 120	Ala	Gly	Ile					
10	(2)	INF	ORMA	SEQU)	FOR ENCE (A) L (B) T	CHA ENGT	RACT H: 1	ERIS 29 a	TICS mino		.ds					
15	Mot	Lvc		SEQ	D) T UENC	OPOL E DE	OGY: SCRI	lin PTIO	ear N: S							
	1		Val	GIŞ	5	Arg	тте	Arg	vai	ьуs 10	Met	Ser	Val	Asn	Lys 15	Al.
20	His	Pro	Val	Val 20	Ser	Thr	His	Trp	Arg 25	Trp	Pro	Ala	Glu	Trp 30	Pro	Gl
	Met	Phe	Leu 35	His	Leu	Ala	Gln	Glu 40	Pro	Arg	Thr	Glu	Val 45	Lys	Ser	Ar
25	Pro	Leu 50	Gly	Leu	Ala	Gly	Phe 55	Ile	Arg	Gln	Asp	Ser 60	Lys	Thr	Arg	Ly
30	Pro 65	Leu	Glu	Gln	Glu	Thr 70	Ile	Met	Ser	Ala	Ala 75	Asp	Thr	Ala	Leu	Trj 80
	Pro	Tyr	Gly	His	Gly 85	Asn	Arg	Glu	His	Gln 90	Glu	Asn	Glu	Leu	Gln 95	Lys
35	Tyr	Leu	Gln	Tyr 100	Lys	Asp	Met	His	Leu 105	Leu	Asp	Ser	Gly	Gln 110	Ser	Let
	Gly	His	Thr 115	His	Thr	Leu	Gln	Gly 120	Ser	His	Asn	Leu	Thr 125	Ala	Leu	Ası
40	Ile															
45	(2)	INF	ORMAI	rion	FOR	SEQ	ID 1	10: 1	L75 :							
50			(i) s	(. (:	A) L B) T D) T	ENGT YPE : OPOL	H: 3 ami: OGY:	72 ai no a lin	mino cid ear	aci		: 17!	5 :			
55	Met 1	Ala	Tyr	His	Ser 5	Phe	Leu	Val	Glu	Pro 10	Ile	Ser	Cys	His	Ala 15	Tr
55	Asn	Lys	Asp	Arg 20	Thr	Gln	Ile	Ala	Ile 25	Cys	Pro	Asn	Asn	His 30	Glu	Va]
60	His	Ile	Tyr 35	Glu	Lys	Ser	Gly	Ala 40	Lys	Trp	Thr	Lys	Val 45	His	Glu	Let

	Lys	Glu 50	His	Asn	Gly	Gln	Val 55	Thr	Gly	Ile	Asp	Trp 60	Ala	Pro	Glu	Ser
5	Asn 65	Arg	Ile	Val	Thr	Cys 70	Gly	Thr	Asp	Arg	Asn 75	Ala	Tyr	Val	Trp	Thr 80
10	Leu	Lys	Gly	Arg	Thr 85	Trp	Lys	Pro	Thr	Leu 90	Val	Ile	Leu	Arg	Ile 95	Asn
10	Arg	Ala	Ala	Arg 100	Cys	Val	Arg	Trp	Ala 105	Pro	Asn	Glu	Asn	Lys 110	Phe	Ala
15	Val	Gly	Ser 115	Gly	Ser	Arg	Val	Ile 120	Ser	Ile	Cys	Tyr	Phe 125	Glu	Gln	Glu
	Asn	Asp 130		·Trp	Val	Cys	Lys 135	His	Ile	Lys	Lys	Pro 140	Ile	Arg	Ser	Thr
20	Val 145	Leu	Ser	Leu	Asp	Trp 150	His	Pro	Asn	Asn	Val 155		Leu	Ala	Ala	Gly 160
25	Ser	Cys	Asp	Phe	Lys 165	Cys	Arg	Ile	Phe	Ser 170		Tyr	Ile	Lys	Glu 175	
25	Glu	. Glu	ı Arg	9 Pro 180		Pro	Thr	Pro	Trp 185		Ser	Lys	Met	Pro 190		Gly
30	Glu	Leu	195		Glu	Ser	Ser	Ser 200		: Cys	; Gly	Trp	Val 205		Gly	Val
	Cys	210		r Ala	Ser	Gly	Ser 215		val	L Ala	a Trp	220		His	: Asp	Ser
35	Th:		l Cy:	s Leu	ı Ala	Asp 230		a Asp	Lys	s Lys	3 Met		a Val	. Ala	a Thr	Leu 240
40	Ala	a Se	r Gl	u Thr	245		Let	ı Lev	ı Ala	a Le		r Phe	e Ile	e Thi	Asr 255	Asn
40	Se	r Le	u Va	1 Ala 260		a Gly	/ His	s Asp	26		e Pr	o Vai	l Leı	270		r Tyr
45	As	p Al	a Al 27		a Gl	y Me	t Le	u Sei 280	_	e Gl	y Gl	y Ar	g Lei 28!		p Val	l Pro
	Ly	s Gl 29		er Se	r Gl	n Ar	g Gl 29		u Th	r Al	a Ar	g Gl 30		g Ph	e Gl:	n Asn
50	Le 30		p Ly	s Ly	s Al	a Se 31		r Gl	u Gl	y Gl	y Th. 31		a Al	a Gl	y Al	a Gly 320
EE	L€	eu As	sp Se	er Le	u Hi 32		s As	n Se	r Va	ıl Se 33		n Il	.e Se	r Va	.1 Le 33	u Ser 5
55	Gl	ly G	ly Ly	ys Al 34		rs Cy	s Se	er Gl	n Ph 34		/s Th	ir Th	ır Gl	у Ме 35		p Gly
60	G]	ly Me		er Il 55	e Tr	p As	sp Vā	1 Ly 36		er Le	eu G∶	lu Se	er Al		eu Ly	s Asp

Leu Lys Ile Lys 370

5

10

(2) INFORMATION FOR SEQ ID NO: 176:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 216 amino acids

- (B) TYPE: amino acid
- (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 176:
- Met Trp Ser Ile Gly Ala Gly Ala Leu Gly Ala Ala Ala Leu Ala Leu 1 5 10 15
- Leu Leu Ala Asn Thr Asp Val Phe Leu Ser Lys Pro Gln Lys Ala Ala 20 25 30
 - Leu Glu Tyr Leu Glu Asp Ile Asp Leu Lys Thr Leu Glu Lys Glu Pro 35 40 45
- Arg Thr Phe Lys Ala Lys Glu Leu Trp Glu Lys Asn Gly Ala Val Ile
 50 55 60
 - Met Ala Val Arg Arg Pro Gly Cys Phe Leu Cys Arg Glu Glu Ala Ala 65 70 75 80
- Asp Leu Ser Ser Leu Lys Ser Met Leu Asp Gln Leu Gly Val Pro Leu 85 90 95
 - Tyr Ala Val Val Lys Glu His Ile Arg Thr Glu Val Lys Asp Phe Gln 100 105 110
- Pro Tyr Phe Lys Gly Glu Ile Phe Leu Asp Glu Lys Lys Phe Tyr
 115 120 125
- Gly Pro Gln Arg Arg Lys Met Met Phe Met Gly Phe Ile Arg Leu Gly 130 135 140
 - Val Trp Tyr Asn Phe Phe Arg Ala Trp Asn Gly Gly Phe Ser Gly Asn 145 150 155 160
- Leu Glu Gly Glu Gly Phe Ile Leu Gly Gly Val Phe Val Val Gly Ser 165 170 175
- Gly Lys Gln Gly Ile Leu Leu Glu His Arg Glu Lys Glu Phe Gly Asp \$180\$ \$185\$ \$190\$
 - Lys Val Asn Leu Leu Ser Val Leu Glu Ala Ala Lys Met Ile Lys Pro 195 200 205
- - (2) INFORMATION FOR SEQ ID NO: 177:

	(i) SEQUENCE CHARACTERISTICS:
	(A) LENGTH: 55 amino acids (B) TYPE: amino acid
	(D) TOPOLOGY: linear
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 177:
	mental mar Cor Val Leu
	Met Lys Pro Val Ser Arg Arg Thr Leu Asp Trp Ile Tyr Ser Val Leu 1 10 15
	1 5 10 13
10	Leu Leu Ala Ile Val Leu Ile Ser Trp Gly Cys Ile Ile Tyr Ala Ser
	20 25 30
	Met Val Ser Ala Arg Arg Gln Leu Arg Lys Lys Tyr Pro Asp Lys Ile
	35 40 45
15	
	Phe Gly Thr Asn Glu Asn Leu 50 55
	50 55
20	TO THE MAN TON GEO IN MO. 179.
	(2) INFORMATION FOR SEQ ID NO: 178:
	(i) SEQUENCE CHARACTERISTICS:
~ ~	(A) LENGTH: 23 amino acids
25	(B) TYPE: amino acid (D) TOPOLOGY: linear
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 178:
20	Met Ala Ala Asn Thr Phe Val Leu Ile Met Gly Ile Pro Thr Ser Ala
30	1 5 10 13
	Asn Ala Xaa Arg Asp Leu Phe
	20
35	
-	
	(2) INFORMATION FOR SEQ ID NO: 179:
	(i) SEQUENCE CHARACTERISTICS:
40	(A) LENGTH: 103 amino acids
	(B) TYPE: amino acid
	(D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 179:
45	Met Ser Ile Cys His Arg Gly Thr Gly Ile Ala Leu Ser Ala Gly Val
	1 5 10 15
	Ser Leu Phe Gly Met Ser Ala Leu Leu Leu Pro Gly Asn Phe Glu Ser
	20 25 30
50	Tyr Leu Glu Leu Val Lys Ser Leu Cys Leu Gly Pro Ala Leu Ile His
	Tyr Leu Giu Leu var Lys Ser Leu Cys Leu Gir 110 1111 201 201 201 201 201 201 201 20
	Thr Ala Lys Phe Ala Leu Val Phe Pro Leu Met Tyr His Thr Trp Asn
55	50 55 60
	Gly Ile Arg His Leu Met Trp Asp Leu Gly Lys Gly Leu Lys Ile Pro
	65 70 75 80
60	Gln Leu Tyr Gln Ser Gly Val Val Val Leu Val Leu Thr Val Leu Ser
50	G1 Due -1

					85					90					95	
5	Ser	Met	Gly	Leu 100	Ala	Ala	Met									
	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO;	180:							
10				(A) I B) T D) T	ENGT YPE : OPOL	H: 4 ami OGY:	8 an no a lir	ear	acid						
15	Met 1	Thr							N: S Pro					Cys	Gln 15	Ile
20	Ser	Gly	Thr	Val 20	Phe	Phe	Phe	Leu	Phe 25	Leu	Phe	Ser	Cys	Phe 30	Leu	Met
25	Gln	Ala	Gln 35	Cys	Asp	Lys	Phe	Val 40	Gly	Trp	Asp	Phe	Phe 45	Phe	Phe	Leu
25																
30	(2)	INF														
35				(. (A) L B) T D) T	ENGT YPE: OPOL	H: 9 ami OGY:	6 am no a lin		acid		: 18	1:			
40	Met 1	Arg	Arg	Ala	Leu 5	Ile	Pro	Pro	Cys	Arg 10	Gly	Gly	Pro	Ser	Ala 15	Ser
				20					Ser 25					30		
45			35					40	Arg				45			
50		50					55		Ala			60				
30	65					70			Pro		75					80
55	₩.A	пåз	GIY	FEO	85	PIO	PIO	Pro	Asp	Pro 90	Pro	Trp	Pro	Val	Thr 95	Leu

	(2)	INFO	RMAT	NOI	FOR	SEQ	ID N	10: 1	.82:							
5			(i) S	(1	A) L: B) T D) T	ENGTI YPE : OPOLO	H: 9: amin DGY:	am: no ao line	ino a cid ear	acid	ı	: 182	2:			
10	Met 1	Leu	Glu	Thr	Thr 5	Lys	His	Val	Gln	Ile 10	Ala	Cys	Met	Leu	Leu 15	Leu
	Thr	Cys	Gln	Ile 20	Phe	Leu	Pro	Ser	Ser 25	Leu	Ser	Pro	Ser	Phe 30	Ile	His
15	Ser	Leu	Thr 35	Asp	Ser	Phe	Ile	Pro 40	Leu	Lys	Lys	Leu	Tyr 45	Val	Cys	Phe
20	Val	Gln 50		Thr	Leu	Leu	Lys 55	Ala	Ala	Gly	Tyr	Lys 60	Ser	Ile	Ser	Glu
20	Ala 65	Leu	Gly	Phe	Asp	Xa a 70	Leu	Leu	Cys	Ser	Ser 75	Ala	Arg	Phe	Val	Trp 80
25	Ile	Cys	His	Thr	Tyr 85	Ser	Arg	Pro	Leu	Val 90		Cys	Ala	Leu	His 95	
30	(2)	INF	ORMA	SEQU	ENCE (A) I	SEQ CHA LENGT TYPE:	RACT TH: 2 : am:	ERIS 27 ar ino a	TICS nino acid		ds					
35			(xi)			CE DE				SEQ I	ID NO): 18	33:			
	Met 1		c Val	. Il∈	e Gly		Lev	ı Leu	i Leu	Val		Ala	ı Lev	ı Gly	Pro 15	Gly
40	Gly	/ Vai	l Sei	Met 20		Glu	ı Lys	. Lys	Lys 25		ı Trş)				
45	(2)	IN	FORM													
50				_	(A) (B) (D)	E CHI LENG TYPE TOPO CE D	TH: : an LOGY	11 a ino : li	mino acid near	aci		0: 1	84:			
55		t Se 1	r Gl	y Gl		u Se: 5	r Ph	e Le	u Le		u Va O	1				
	(2) IN	FORM	OITA	N FO	R SE	Q ID	NO:	185	:						
60			(3)	SEC.	או וביאור	יד כו	מפאנו	יייבים ו	STIC	·s.						

```
(A) LENGTH: 65 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 185:
 5
      Met Phe Ala Asp Phe Ile Val Val Thr Ala Thr Val Gln Arg Cys Pro
      Gly Ser Pro Pro Leu Ser Glu Ile Leu Trp Lys Asp Glu Pro Phe Ala
10
                   20
                                       25
      Ile Ser Ser His Ala Gly Leu Pro Trp Leu Ser Ser Trp Pro Ala Pro
                                   40
15
      Pro Trp Thr Trp Ser Trp Ile Ser Arg Arg Arg Glu His Gly Arg Gly
      Ser
       65
20
      (2) INFORMATION FOR SEQ ID NO: 186:
25
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 22 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 186:
30
      Met Val Glu Ser Val Met Pro Val Val Val Cys Thr Leu Ser Pro Gly
                                           1.0
      Ile Asp Ser Ser Pro Ser
35
                   20
      (2) INFORMATION FOR SEQ ID NO: 187:
40
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 132 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
45
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 187:
      Met Asp Val Leu Phe Val Ala Ile Phe Ala Val Pro Leu Ile Leu Gly
50
      Gln Glu Tyr Glu Asp Glu Glu Arg Leu Gly Glu Asp Glu Tyr Tyr Gln
                   20
                                       25
      Val Val Tyr Tyr Tyr Thr Val Thr Pro Ser Tyr Asp Asp Phe Ser Ala
55
      Asp Phe Thr Ile Asp Tyr Ser Ile Phe Glu Ser Glu Asp Arg Leu Asn
      Arg Leu Asp Lys Asp Ile Thr Glu Ala Ile Glu Thr Thr Ile Ser Leu
60
                           70
                                               75
```

	Glu	Thr	Ala	Arg	Ala 85	Asp	His	Pro	Lys	Pro 90	Val	Thr	Val	Lys	Pro 95	Val
5	Thr	Thr	Glu	Pro 100	Gln	Ser	Pro	Asp	Leu 105	Asn	Asp	Ala	Val	Ser 110	Ser	Leu
10	Arg	Ser	Pro 115	Ile	Pro	Leu	Leu	Leu 120	Ser	Cys	Ala	Phe	Val 125	Gln	Val	Gly
10	Met	Tyr 130	Phe	Met												
15	(2)	INF	· ORMA	rion	FOR	SEQ	ID	NO:	188:							
20				(A) L B) T D) T	ENGT YPE : 'OPOL	H: 6 ami OGY:	9 am no a lin		acid		: 18	8:			
25	Met 1	Pro	Суз	Gln	Pro 5	Gly	Gln	Val	Pro	Ser 10	Cys	Gln	Cys	Thr	Phe 15	
	Leu	Leu	Leu	Met 20	Leu	Pro	Ser	Leu	Pro 25	Ser	Pro	Ala	Ser	Gln 30		Arg
30	Pro	Phe	Суs 35		Ser	Met	Glu	Tyr 40	Phe	His	Gly	Cys	Ala 45	Ser	Pro	Ser
35	Gln	Ala 50		Ile	Gly	Gly	Phe 55		Phe	Ala	Ser	Val 60		Leu	Ala	Asp
,	Ile 65		. Cys	Leu	Gln											
40	(2)	INF	ORMA	TION	FOR	SEQ] ID	NO:	189:							
45				~	(A) 1 (B) '	LENG: FYPE FOPOI	TH: 4 : am: LOGY	45 ar ino a : lir		acio		D: 1 8	39:			
50	Met		. Leu	ı Lev	Ser 5		Ala	ı Ile	e Pro	Ala 10		ı Thr	: Leu	ı Ile	e Phe	
	Leu	ı Met	Phe	Phe		Phe	e Pro) Phe	e Arg		a His	s Thi	C Val	. Val		r Ile
55	Va]	l Alá	a Ser 35	_	, Phe	e Lei	ı Gly	/ Let 4(ı Ser	r Pro	Let	ı Cys	s Gl _y 45			
60	(2)) INI	FORM	OITA	ı FOI	R SE(Q ID	NO:	190	:						

5				(A) L B) T D) T	ENGT: YPE: OPOL	H: 6 ami OGY:	5 am no a lin	ino d cid ear	acid		. 10	0			
10	Met 1													Pro	Туг 15	Pro
10	Leu	Gln	Trp	Ser 20	Leu	Leu	Val	Ala	Val 25	Val	Ala	Gly	Ser	Val 30	Val	Ser
15	Tyr	Gly	Val 35	Thr	Arg	Val	Glu	Ser 40	Glu	Lys	Cys	Asn	Asn 45	Leu	Trp	Leu
	Phe	Leu 50	Glu	Thr	Gly	Gln	Leu 55	Pro	Lys	Asp	Arg	Ser 60	Thr	Asp	Gln	Arg
20	Ser 65															
25	(2)	INF	ORMA'	rion	FOR	SEQ	ID 1	NO: 1	L91:							
30				(A) L B) T D) T	ENGT YPE: OPOL	H: 5 ami OGY:	0 am no a lin	ino cid ear	acid		: 19	1:			
35	Met 1	Asn	Leu	Leu	Gly 5	Met	Ile	Phe	Ser	Met 10	Cys	Gly	Leu	Met	Leu 15	Lys
	Leu	Lys	Trp	Cys 20	Ala	Trp	Val	Ala	Val 25	Tyr	Cys	Ser	Phe	Ile 30	Ser	Phe
40	Ala	Asn	Ser 35	Arg	Ser	Ser	Glu	Asp 40	Thr	Lys	Gln	Met	Met 45	Ser	Ser	Phe
	Met	Xaa 50														
45	(2)	TNE	ODMAI	TION	EOD	CEO.	TD.	NO.	100.							
50	(2)	1141.	(i)	SEQU (ENCE A) L B) T D) T	CHA ENGT YPE:	RACT H: 1 ami OGY:	ERIS 70 a no a lin	TICS mino cid ear	aci		: 19	2 :			
55	Met 1		Leu	Asn	Val 5	Ala	Leu	Val	Ala	Leu 10	Val	Leu	Leu	Gly	Ala 15	Tyr
60	Arg	Leu	Trp	Val 20		Trp	Gly	Arg	Arg 25		Leu	Gly	Ala	Gly 30	Ala	Gly

	Ala	Gly	Glu 35	Glu	Ser	Pro	Ala	Thr 40	Ser	Leu	Pro	Arg	Met 45	Lys	Lys	Arg
5	Asp	Phe 50	Ser	Leu	Glu	Gln	Leu 55	Arg	Gln	Tyr	Asp	Gly 60	Ser	Arg	Asn	Pro
	Arg 65	Ile	Leu	Leu	Ala	Val 70	Asn	Gly	Lys	Val	Phe 75	Asp	Val	Thr	Lys	Gly 80
10	Ser	Lys	Phe	Tyr	Gly 85	Pro	Ala	Gly	Pro	Туг 90	Gly	Ile	Phe	Ala	Gly 95	Arg
15	Asp	Ala	Ser	Arg 100	Gly	Leu	Ala	Thr	Phe 105	Cys	Leu	Asp	Lys	Asp 110	Ala	Leu
13	Arg	Asp	Glu 115		Asp	Asp	Leu	Ser 120	Asp	Leu	Asn	Ala	Val 125	Gln	Met	Glu
20	Ser	Val 130	Arg	Glu	Trp	Glu	Met 135		Phe	Lys	Glu	Lys 140		Asp	Tyr	Val
	Gly 145		Leu	Leu	Lys	Pro 150	Gly	Glu	Glu	Pro	Ser 155		Tyr	Thr	Asp	Glu 160
25	Glu	Asp	Thr	Lys	Asp 165		Asn	. Lys	Gln	Asp 170						
30	(2)	INF		ATION SEQU												
35					(A) : (B) ' (D) '	LENG: TYPE TOPO!	TH: : am LOGY	66 an ino a : lin	mino acid near	aci		D: 1	93:			
40	Met 1		тул	r Phe		c Gly	/ Let	ı Leı	ı Val	1 Ile 10		ı Ala	a Phe	≥ Alá	a Ala	a Trp
40	Va]	L Ala	a Le	u Ala 20		ı Gly	, Lei	ı Gly	y Val 25		a Va	l Ty:	r Ala	a Ala 30		a Val
45	Let	ı Le	u Gl; 3		a Gl	у Суз	s Ala	a Thi		e Lei	ı Va	l Th	r Se:		ı Ala	a Met
	Thi	r Ala		p Le	u Il	e Gly	y Pro 5		s Th	r Ası	n Se	r Gl		u Se:	r Cy:	s Thr
50	Ala 6!	a Pro	0													
55	(2) IN	FORM	ATIO	N FO	R SE	Q ID	NO:	194	:						
			(i)	SEC	(A)	E CH LENC TYPE	TH:	92 a	amino	aci	ids					
60						TOPO										

			(xi)	SEQ	UENC	E DE	SCRI	PTIO	N: S	EQ I	D NO	: 19	4:			
5	Met 1	Ala	Ala	Gly	Pro 5	Ser	Gly	Cys	Leu	Val 10	Pro	Ala	Phe	Gly	Leu 15	Arg
	Leu	Leu	Leu	Ala 20	Thr	Val	Leu	Gln	Ala 25	Val	Ser	Ala	Phe	Gly 30	Ala	Glu
10	Phe	Ser	Ser 35	Glu	Ala	Cys	Arg	Glu 40	Leu	Gly	Phe	Ser	Ser 45	Asn	Leu	Leu
	Cys	Ser 50	Ser	Cys	Asp	Leu	Leu 55	Gly	Gln	Phe	Asn	Leu 60	Leu	Gln	Leu	Asp
15	Pro 65	Asp	Cys	Arg	Gly	Cys 70	Cys	Gln	Glu	Glu	Ala 75	Gln	Phe	Glu	Thr	Lys 80
20	Lys	Leu	Tyr	Ala	Gly 85	Ala	Ile	Leu	Glu	Val 90	Cys	Gly				
	(2)		ORMAT													
25			(i) :	(A) L B) T	CHAI ENGT YPE: OPOL	H: 1 ami	76 a no a	mino cid		ds					
30			(xi)							EQ II	D NO	: 19	5:			
	Met 1	Arg	Gly	Ser	His 5	Leu	Arg	Leu	Leu	Pro 10	Tyr	Leu	Val	Ala	Ala 15	Asn
35	Pro	Val	Asn	Tyr 20	Gly	Arg	Pro	Tyr	Arg 25	Leu	Ser	Cys	Val	Glu 30	Ala	Phe
	Ala	Ala	Thr 35	Phe	Cys	Ile	Val	Gly 40	Phe	Pro	Asp	Leu	Ala 45	Val	Ile	Leu
40	Leu	Arg 50	Lys	Phe	Lys	Trp	Gly 55	Lys	Gly	Phe	Leu	Asp 60	Leu	Asn	Arg	Gln
45	65					70					75				Leu	80
					85					90					Glu 95	
50				100					105					110	Asn	
	Asn	Arg	Pro 115	Val	Ala	Ser	Thr	Arg 120	Leu	Pro	Ser	Asp	Thr 125	Asp	Asp	Ser
55		130					135					140			Ala	
60	Ser 145	Ser	Cys	Cys	Glu	Glu 150	Glu	Gln	Thr	Gln	Gly 155	Arg	Gly	Ala	Glu	Ala 160

Arg Ala Pro Ala Glu Val Trp Lys Gly Ile Lys Lys Arg Gln Arg Asp 170 5 (2) INFORMATION FOR SEQ ID NO: 196: 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 70 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 196: 15 Met Ser Asn Ala Cys Lys Glu Leu Ala Ile Phe Leu Thr Thr Gly Ile 10 5 Val Val Ser Ala Phe Gly Leu Pro Ile Val Phe Ala Arg Ala His Leu 20 Ile Glu Trp Gly Ala Cys Ala Leu Val Leu Thr Gly Asn Thr Val Ile 40 25 Phe Ala Thr Ile Leu Gly Phe Phe Leu Val Phe Gly Ser Asn Asp Asp 55 Phe Ser Trp Gln Gln Trp 30 65 (2) INFORMATION FOR SEQ ID NO: 197: 35 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 25 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear 40 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 197: Met Thr Leu Leu Ile Ile Phe Leu Pro Phe Xaa Phe Thr Thr Xaa Thr 10 1 5 45 Asn Ser Gly Gly Ser Phe Pro Val Arg 20 (2) INFORMATION FOR SEQ ID NO: 198: 50 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 73 amino acids (B) TYPE: amino acid 55 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 198: Met Lys Gly Glu Leu Leu Pro Phe Leu Phe Leu Thr Val Trp Leu Trp 1 5 60

	Leu	Tyr	Lys	Leu 20	Xaa	Phe	Gly	Glu	Ser 25	Pro	Arg	Tyr	Pro	Asn 30	Val	Ile
5	Gly	Lys	Thr 35	Tyr	Phe	Phe	Phe	Trp 40	Thr	Asp	Gln	Ile	Ser 45	Arg	Glu	Ser
	Arg	Phe 50	Leu	Glu	Arg	Leu	Ala 55	Phe	Ile	Val	Ser	Glu 60	Asn	Cys	Leu	Ile
10	Phe 65	Leu	Ile	His	Ala	Ile 70	Thr	Gly	Gln							
15	(2)	INF	ORMA													
20			(i) (xi)	(A) L B) T D) T	CHA: ENGT YPE: OPOL E DE:	H: 2 ami OGY:	89 a no a lin	mino cid ear	aci		: 19	9:			
25	Met 1	Ser	Gly	Phe	Ser 5	Thr	Glu	Glu	Arg	Ala 10	Ala	Pro	Phe	Ser	Leu 15	Glu
23	Tyr	Arg	Val	Phe 20	Leu	Lys	Asn	Glu	Lys 25	Gly	Gln	Tyr	Ile	Ser 30	Pro	Phe
30	His	Asp	Ile 35	Pro	Ile	Tyr	Ala	Asp 40	Lys	Asp	Val	Phe	His 45	Met	Val	Val
	Glu	Val 50	Pro	Arg	Trp	Ser	Asn 55	Ala	Lys	Met	Glu	Ile 60	Ala	Thr	Lys	Asp
35	Pro 65	Leu	Asn	Pro	Ile	Lys 70	Gln	Asp	Val	Lys	Lys 75	Gly	Lys	Leu	Arg	Туr 80
40	Val	Ala	Asn	Leu	Phe 85	Pro	Tyr	Lys	Gly	Туг 90	Ile	Trp	Asn	Tyr	Gly 95	Ala
	Ile	Pro	Gln	Thr 100	Trp	Glu	Asp	Pro	Gly 105	His	Asn	Asp	Lys	His 110	Thr	Gly
45	Cys	Суѕ	Gly 115	Asp	Asn	Asp	Pro	Ile 120	Asp	Val	Cys	Glu	Ile 125	Gly	Ser	Lys
	Val	Cys 130	Ala	Arg	Gly	Glu	Ile 135	Ile	Gly	Val	Lys	Val 140	Leu	Gly	Ile	Leu
50	Ala 145	Met	Ile	Asp	Glu	Gly 150	Glu	Thr	Asp	Trp	Lys 155	Val	Ile	Ala	Ile	Asn 160
55	Val	qzA	Asp	Pro	Asp 165	Ala	Ala	Asn	Tyr	Asn 170	Asp	Ile	Asn	Asp	Val 175	Lys
	Arg	Leu	Lys	Pro 180	Gly	Tyr	Leu	Glu	Ala 185	Thr	Val	Asp	Trp	Phe 190	Arg	Arg
60	Tyr	Lys	Val 195	Pro	Asp	Gly	Lys	Pro 200	Glu	Asn	Glu	Phe	Ala 205	Phe	Asn	Ala

	Glu	Phe 210	Lys	Asp	Lys		Pne . 215	Ala	TTE	ASP	116	220	БУБ	ser	1111	*****
5	Asp 225	His	Trp	Lys	Ala	Leu 230	Val	Thr	Lys	Lys	Thr 235	Asn	Gly	Lys	Gly	Ile 240
10	Ser	Cys	Met	Asn	Thr 245	Thr	Leu	Ser	Glu	Ser 250	Pro	Phe	Lys	Cys	Asp 255	Pro
(U	Asp	Ala	Ala	Arg 260	Ala	Ile	Val	Asp	Ala 265	Leu	Pro	Pro	Pro	Cys 270	Glu	Ser
15	Ala	Cys	Thr 275	Val	Pro	Thr	Asp	Val 280	Asp	Lys	Trp	Phe	His 285		Gln	Lys
	Asn															
20																,
	(2)	INF		TION												
25			(i)		(A) I	: CHA LENGT TYPE:	H: 6	25 a	amino		ids					
			(xi) SEÇ		CE DE				SEQ I	ED NO	D: 20	00:			
30	Met 1		ı Ile	e Pro	Gly 5		Leu	Cys	: Lys	Lys 10		L Lys	s Lev	ı Ser	Asn 15	Asn
35	Ala	Gl:	n Ası	n Trp 20		Met	Gln	. Arg	y Ala 25		: Ası	n Val	l Thi	с Туз 30		n Ala
33	His	Hi	s Va 3		c Arg	g Asn	Lys	Arg 40		/ Gli	n Va	l Va	1 Gly 49		r Arg	g Gly
40,	Gly		e Ar O	g Gly	y Cys	s Thr	Val		o Lei	ı Thi	r Gl	y Le		r Gl	y Ala	a Gly
	Ly: 6!		r Th	r Va	l Sei	r Met 70		a Le	u Gl	u Gl	и Ту 7		u Va	l Cy	s Hi	s Gly 80
45	Il	e Pr	о Су	rs Ty	r Thi		ı Asp	o Gl	y As		n Il O	e Ar	g Gl	n Gl	y Le	u As n 5
50	Ly	s As	n L∈	u Gl 10		e Se:	r Pro	o Gl	u As 10		g Gl	u Gl	u As	n Va 11		g Arg
50	11	e Al		lu Va L5	.1 Al	a Ly	s Le	u Ph 12		a As	p Al	a Gl	.у Le 12		ıl Cy	rs Ile
55	Th		er Pl 30	ne Il	e Se	er Pr	о Ту 13		ır Gl	.n As	ıA qı		sn As 10	sn Al	.a Ar	g Gln
	11 14		is G	lu Gl	ly Al	.a Se 15	_	u Pr	co Ph	ne Ph		lu Va 55	al Ph	ne Va	al As	p Ala 160
60	Pı	o L	eu H	is Va	al Cy	ys Gl	u Gl	n Aı	rg As	sp Va	al L	ys G	ly L	eu T	yr Ly	ys Lys

					165					170					175	
5	Ala	Arg	Ala	Gly 180	Glu	Ile	Lys	Gly	Phe 185		Gly	Ile	Asp	Ser 190		Tyr
	Glu	Lys	Pro 195	Glu	Ala	Pro	Glu	Leu 200	Val	Leu	Lys	Thr	Asp 205		Cys	Asp
10	Val	Asn 210	Asp	Cys	Val	Gln	Gln 215		Val	Glu	Leu	Leu 220	Gln	Glu	Arg	Asp
	Ile 225	Val	Pro	Val	Asp	Ala 230	Ser	Tyr	Glu	Val	Lys 235	Glu	Leu	Tyr	Val	Pro 240
15	Glu	Asn	Lys	Leu	His 245	Leu	Ala	Lys	Thr	Asp 250	Ala	Glu	Thr	Leu	Pro 255	Ala
20	Leu	Lys	Ile	Asn 260	Lys	Val	Asp	Met	Gln 265	Trp	Val	Gln	Val	Leu 270	Ala	Glu
	Gly	Trp	Ala 275	Thr	Pro	Leu	Asn	Gly 280	Phe	Met	Arg	Glu	Arg 285	Glu	Tyr	Leu
25	Gln	Cys 290	Leu	His	Phe	Asp	Cys 295	Leu	Leu	Asp	Gly	Gly 300	Val	Ile	Asn	Leu
	Ser 305	Val	Pro	Ile	Val	Leu 310	Thr	Ala	Thr	His	Glu 315	Asp	Lys	Glu	Arg	Leu 320
30	Asp	Gly	Cys	Thr	Ala 325	Phe	Ala	Leu	Met	Tyr 330	Glu	Gly	Arg	Arg	Val 335	Ala
35				340				Phe	345					350		
			355					Cys 360					365			
40		370					375	Leu				380				
	Asp 385	Arg	Val	Tyr	Trp	Asn 390	Asp	Gly	Leu	Asp	Gln 395	Tyr	Arg	Leu	Thr	Pro 400
45	Thr	Glu	Leu	Lys	Gln 405	Lys	Phe	Lys	Asp	Met 410	Asn	Ala	Asp	Ala	Val 415	Phe
50	Ala	Phe	Gln	Leu 420	Arg	Asn	Pro	Val	His 425	Asn	Gly	His	Ala	Leu 430	Leu	Met
	Gln	Asp	Thr 435	His	Lys	Gln	Leu	Leu 440	Glu	Arg	Gly	Tyr	Arg 445	Arg	Pro	Val
55	Leu	Leu 450	Leu	His	Pro	Leu	Gly 455	Gly	Trp	Thr	Lys	Asp 460	Asp	Asp	Val	Pro
	Leu 465	Met	Trp	Arg	Met	Lys 470	Gln	His	Ala	Ala	Val 475	Leu	Glu	Glu	Gly	Val 480
60	Leu	Asn	Pro	Glu	Thr	Thr	Val	Val	Ala	Ile	Phe	Pro	Ser	Pro	Met	Met

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					485					490					495	
E	Tyr	Ala	Gly	Pro 500	Thr	Glu	Val	Gln	Trp 505	His	Cys	Arg	Ala	Arg 510	Met	Val
5	Ala	Gly	Ala 515	Asn	Phe	Tyr	Ile	Val 520	Gly	Arg	Asp	Pro	Ala 525	Gly	Met	Pro
10	His	Pro 530	Glu	Thr	Gly	Lys	Asp 535	Leu	Тут	Glu	Pro	Ser 540	His	Gly	Ala	Lys
	Val 545	Leu	Thr	Met	Ala	Pro 550	Gly	Leu	Ile	Thr	Leu 555	Glu	Ile	Val	Pro	Phe 560
15	Arg	Val	Ala	Ala	Туr 565	Asn	Lys	Lys	Lys	Lys 570		Met	Asp	Tyr	тут 575	Asp
20	Ser	Glu	His	His 580	Glu	Asp	Phe	Glu	Phe 585		Ser	Gly	Thr	Arg 590	Met	Arg
20	Lys	Leu	Ala 595	Arg	Glu	Gly	Gln	Lys 600		Pro	Glu	Gly	Phe 605		Ala	Pro
25	Lys	Ala 610		Thr	Val	Leu	Thr 615		Tyr	Tyr	Lys	Ser 620	Leu	Glu	Lys	Ala
	Xaa 625															
30																
	(2)	INF	ORMA	TION	FOF	SEÇ) ID	NO:	201:	:						
35					(A) : (B) ' (D) '	LENG TYPE TOPO	TH: : am LOGY	649 ino : li	STIC: amin acid near	o ac			0.1			
40									ON:					- D~	. T.17	- Pro
40		: Se: L	r Ala	a Sei		n Asj	o re	ı Gi	u Pro	о гу: 1) Le	1 211	e PIC	19 19	Pro
45	Ala	a Ph	e Gly						u Se		r Gli	u Ası	n Se	r His		ı Asp
43	Glı	ı Se	r Pro		t Ly:	s As	n Va	1 Se 4		r Se	r Ly	s Gl	y Se 4		o Al	a Pro
50	Le		y Va O	l Ar	g Se	r Ly		r Gl 5	y Pr	o Le	u Ly	s Pr 6		a Ar	g Gl	u Asp
		r Gl 5	u As	n Ly	s As		s Al	a Gl	y Gl	u Il		r Se 5	r Le	u Pr	o Ph	e Pro 80
55	Gl	y Va	ıl Va	l Le		s Pr	o Al	a Al	a Se		g G1 90	y Gl	y Pr	o Gl		u Sei 5
60	Ly	s As	sn Gl	y Gl 10		u Ly	/s Ly	rs Gl	lu As 10		rg Ly	s Il	e As	sp Al 11		a Ly

	Asr	n Thi	r Phe 119	e Glr	ı Ser	. Lys	Il€	Asr 120	ı Glr	ı Glu	ı Glu	ı Let	125		Gly	/ Thr
5	Pro	Pro 130	o Ala	a Arg	y Ph∈	Pro	Lys 135	Ala	Pro	Ser	Lys	Leu 140		· Val	l Gly	/ Gly
	Pro 145	Tr	Gly	/ Glr	ser	Gln 150	Glu	Lys	Glu	Lys	Gly 155		Lys	s Asr	ser	Ala 160
10	Thr	Pro) Lys	Gln	Lys 165	Pro	Leu	Pro	Pro	Leu 170		Thr	Leu	ı Gly	Pro 175	Pro
15	Pro	Pro	Lys	Pro 180	Asn	Arg	Pro	Pro	Asn 185		. Asp	Leu	Thr	Lys 190		His
	Lys	Thr	Ser 195	Ser	Gly	Asn	Ser	Thr 200	Ser	Lys	Gly	Gln	Thr 205		Tyr	Ser
20	Thr	Thr 210	Ser	Leu	Pro	Pro	Pro 215	Pro	Pro	Ser	His	Pro 220		. Ser	Gln	. Pro
	Pro 225	Leu	Pro	Ala	Ser	His 230	Pro	Ser	Gln	Pro	Pro 235	Val	Pro	Ser	Leu	Pro 240
25	Pro	Arg	Asn	Ile	Lys 245	Pro	Pro	Phe	Asp	Leu 250	Lys	Ser	Pro	Val	Asn 255	Glu
30	Asp	Asn	Gln	Asp 260	Gly	Val	Thr	His	Ser 265	Asp	Gly	Ala	Gly	Asn 270	Leu	Asp
	Glu	Glu	Gln 275	Asp	Ser	Glu	Gly	Glu 280	Thr	Tyr	Glu	Asp	Ile 285	Glu	Ala	Ser
35	Lys	Glu 290	Arg	Glu	Lys	Lys	Arg 295	Glu	Lys	Glu	Glu	Lys 300	Lys	Arg	Leu	Glu
	Leu 305	Glu	Lys	Lys•	Glu	Gln 310	Lys	Glu	Lys	Glu	Lys 315	Lys	Glu	Gln	Glu	Ile 320
40	Lys	Lys	Lys	Phe	Lys 325	Leu	Thr	Gly	Pro	Ile 330	Gln	Val	Ile	His	Leu 335	Ala
45	Lys	Ala	Cys	Cys 340	Asp	Val	Lys	Gly	Gly 345	Lys	Asn	Glu	Leu	Ser 350	Phe	Lys
	Gln	Gly	Glu 355	Gln	Ile	Glu	Ile	Ile 360	Arg	Ile	Thr	Asp	Asn 365	Pro	Glu	Gly
50	Lys	Trp 370	Leu	Gly	Arg	Thr	Ala 375	Arg	Gly	Ser	Tyr	Gly 380	Tyr	Ile	Lys	Thr
	Thr 385	Ala	Val	Glu	Ile	Asp 390	Tyr	Asp	Ser	Leu	Lys 395	Leu	Lys	Lys	Asp	Ser 400
55	Leu	Gly	Ala	Pro	Ser 405	Arg	Pro	Ile		Asp 410	Asp	Gln	Glu	Val	Tyr 415	Asp
50	Asp	Val	Ala	Glu 420	Gln	Asp	Asp	Ile	Ser 425	Ser	His	Ser	Gln	Ser 430	Gly	Ser

	Gly	Gly	11e 435	Phe	Pro	Pro	Pro	Pro 440	Asp	Asp	Asp	Ile	Туг 445	Asp	Gly	Ile
5	Glu	Glu 450	Glu	Asp	Ala	Asp	Asp 455	Gly	Ser	Thr	Leu	Gln 460	Val	Gln	Glu	Lys
	Ser 465	Asn	Thr	Trp	Ser	Trp 470	Gly	Ile	Leu	Lys	Met 475	Leu	Lys	Gly	Lys	Asp 480
10	Asp	Arg	Lys	Lys	Ser 485	Ile	Arg	Glu	Lys	Pro 490	Lys	Val	Ser	Asp	Ser 495	Asp
15	Asn	Asn	Glu	Gly 500	Ser	Ser	Phe	Pro	Ala 505	Pro	Pro	Lys	Gln	Leu 510	Asp	Met
13	Gly	Asp	Glu 515	Val	Tyr	Asp	Asp	Val 520	Asp	Thr	Ser	Asp	Phe 525	Pro	Val	Ser
20	Ser	Ala 530	Glu	Met	Ser	Gln	Gly 535	Thr	Asn	Val	Gly	Lys 540	Ala	Lys	Thr	Glu
	Glu 545	Lys	Asp	Leu	Lys	Lys 550	Leu	Lys	Lys	Gln	Xaa 555		Xaa	Xaa	Lys	Asp 560
25	Phe	Arg	Lys	Lys	Phe 565	Lys	Tyr	Asp	Gly	Glu 570		Arg	Val	Leu	Tyr 575	Ser
30	Thr	Lys	Val	Thr 580	Thr	Ser	Ile	Thr	Ser 585		Lys	Trp	Gly	Thr 590	Arg	Asp
	Leu	Gln	Val 595		Pro	Gly	Glu	Ser 600		Glu	ı Val	Ile	605		Thr	Asp
35	Asp	Thr 610		Val	Leu	Cys	Arg 615		Glu	. Glu	ı Gly	620		· Gly	Tyr	Val
	Leu 625	_	ßer	Туг	Leu	Ala 630) Asn	Asp	Gly	7 Glu 635		туг	: Asp	Asp	11e 640
40	Ala	Asp	Gly	Cys	11e 645		Asp	Asn	Asp)						
45	(2)	INE			1 FOR											
50					UENCE (A) I (B) ' (D) '	LENG IYPE IOPO	TH: : am LOGY	55 a ino : li	mino acid near	aci		0: 2	02:			
<i></i>		t Ala	a Trj	o Pro	ser		g Sei	r Lys	s Met	t Ph 1		r Le	u Le	u Pro	o Va:	Leu 5
55	Суя	з Ту	r Le	u Trj 2		r Le	u Trj	o Le	u Pro 2!		n Ph	e Se	r Tr	p Ile 3		n Glu
60	Le	u Ly	s Al		l Le	ı Ar	g As	p As		y Le	u Il	e Se	r Al 4		l Al	a Trp

	Asn	Ala 50	Glu	Phe	Gln	Thr	Cys 55									
5																
	(2)	INF	ORMA	TION	FOR	SEQ	ID :	NO:	203:							
10				(((A) I (B) T (D) T	ENGT YPE : OPOL	RACT H: 2 ami OGY: SCRI	67 a no a lin	mino cid ear	aci		: 20	3:			
15	Met 1	Val	Lys	Val	Thr 5	Phe	Asn	Ser	Ala	Leu 10	Ala	Gln	Lys	Glu	Ala 15	Lys
20	Lys	Asp	Glu	Pro 20	Lys	Ser	Gly	Glu	Glu 25	Ala	Leu	Ile	Ile	Pro 30	Pro	Asp
	Ala	Val	Ala 35	Val	Asp	Cys	Lys	Asp 40	Pro	Asp	Asp	Val	Val 45	Pro	Val	Gly
25	Gln	Arg 50	Arg	Ala	Trp	Cys	Trp 55	Cys	Met	Cys	Phe	Gly 60	Leu	Ala	Phe	Met
	Leu 65	Ala	Gly	Val	Ile	Leu 70	Gly	Gly	Ala	Tyr	Leu 75	Tyr	Lys	Tyr	Phe	Ala 80
30	Leu	Gln	Pro	Asp	Asp 85	Val	Tyr	Tyr	Cys	Gly 90	Ile	Lys	Tyr	Ile	Lys 95	Asp
35	Asp	Val	Ile	Leu 100	Asn	Glu	Pro	Ser	Ala 105	Asp	Ala	Pro	Ala	Ala 110	Leu	Tyr
	Gln	Thr	Ile 115	Glu	Glu	Asn	Ile	Lys 120	Ile	Phe	Glu	Glu	Glu 125	Glu	Val	Glu
40	Phe	Ile 130	Ser	Val	Pro	Val	Pro 135	Glu	Phe	Ala	Asp	Ser 140	Asp	Pro	Ala	Asn
	Ile 145	Val	His	Asp	Phe	Asn 150	Lys	Lys	Leu	Thr	Ala 155	Tyr	Leu	Asp	Leu	Asn 160
45	Leu	Asp	Lys	Cys	Tyr 165	Val	Ile	Pro	Leu	Asn 170	Thr	Ser	Ile	Val	Met 175	Pro
50	Pro	Arg	Asn	Leu 180	Leu	Glu	Leu	Leu	Ile 185	Asn	Ile	Lys	Ala	Gly 190	Thr	Tyr
	Leu	Pro	Gln 195	Ser	Tyr	Leu	Ile	His 200	Glu	His	Met	Val	Ile 205	Thr	Asp	Arg
55	Ile	Glu 210	Asn	Ile	Asp	His	Leu 215	Gly	Phe	Phe	Ile	Tyr 220	Arg	Leu	Cys	His
	Asp 225	Lys	Glu	Thr	Tyr	Lys 230	Leu	Gln	Arg	Arg	Glu 235	Thr	Ile	Lys	Gly	Ile 240
60	Gln	Lys	Arg	Glu	Ala	Ser	Asn	Cys	Phe	Ala	Ile	Arg	His	Phe	Glu	Asn

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				2	245				2	250				:	2 5 5	
5	Lys I	Phe .		Val (260	Glu 1	Thr I	Leu		Cys 8 265	Ser 2	Kaa					
10	(2)							O: 2								
10				(P (E) LE 3) TY)) TO	NGTH PE: POLC	H: 31 amir XGY:	l5 am no ac line	ino id ar			204	:			
15	Met			_				Met						Thr	Ala 15	Phe
20	Ala	Leu	Ser	Lys 20	Pro	Thr	Glu	Lys	Lys 25	Asp	Arg	Val :	His	His 30	Glu	Pro
	Gln	Leu	Ser 35	Asp	Lys	Val	His	Asn 40	Asp	Ala	Gln	Ser	Phe 45	Asp	Tyr	Asp
25	His	Asp 50	Ala	Phe	Leu	Gly	Ala 55	Glu	Glu	Ala	Lys	Thr 60	Phe	Asp	Gln	Leu
30	Thr 65	Pro	Glu	Glu	Ser	Lys 70	Glu	Arg	Leu	Gly	Lys 75	Ile	Val	Ser	Lys	Ile 80
50	Asp	Gly	Asp	Lys	Asp 85	Gly	Phe	Val	Thr	Val 90	Asp	Glu	Leu	Lys	Asp 95	Trp
35	Ile	Lys	Phe	Ala 100	Gln	Lys	Arg	Trp	Ile 105	Tyr	Glu	Asp	Val	Glu 110	Arg	Gln
	Trp	Lys	Gly 115	His	Asp	Leu	Asn	Glu 120	Asp	Gly	Leu	Val	Ser 125	Trp	Glu	Glu
40	Tyr	Lys 130		Ala	Thr	Tyr	Gly 135	Tyr	Val	Leu	Asp	Asp 140	Pro	Asp	Pro	Asp
45	Asp 145		Phe	Asn	Tyr	Lys 150	Gln	Met	Met	Val	Arg 155	Asp	Glu	Arg	Arg	Phe 160
	Lys	Met	Ala	Asp	Lys 165	Asp	Gly	Asp	Leu	Ile 170	Ala	Thr	Lys	Glu	Glu 175	Phe
50	Thr	Ala	Phe	Leu 180		Pro	Glu	Glu	Tyr 185		Tyr	Met	Lys	Asp 190	Ile	Val
	Val	. Glr	195		Met	Glu	Asp	200		Lys	Asn	Ala	Asp 205		Phe	Ile
55	Asp	210		ı Glu	Тух	Ile	215	/ Asp	Met	Туг	Ser	His 220		Gly	Asn	Thr
60	Asp 225		ı Pro	Glu	Trp	230		s Thr	Glu	Arg	235		Phe	e Val	. Glu	Phe 240

	Arg	Asp	Lys	Asn	Arg 245	qzA	Gly	Lys	Met	Asp 250	Lys	Glu	Glu	Thr	Lys 255	Asp
5	Trp	Ile	Leu	Pro 260	Ser	Asp	Tyr	Asp	His 265	Ala	Glu	Ala	Glu	Ala 270	Arg	His
	Leu	Val	Тут 275	Glu	Ser	Asp	Gln	Asn 280	Lys	Asp	Gly	Lys	Leu 285	Thr	Lys	Glu
10	Glu	Ile 290	Val	Asp	Lys	Tyr	Asp 295	Leu	Phe	Val	Gly	Ser 300	Gln	Ala	Thr	Asp
15	Phe 305	Gly	Glu	Ala	Leu	Val 310	Arg	His	Asp	Glu	Phe 315					
20	(2)		ORMA:	SEQUI () ()	ENCE A) L B) T	CHAI ENGT	RACT H: 2 ami	ERI <i>s</i> 07 a no a	rics mino cid		ds					
25	Met 1		(xi) Asp											Lys	Asp 15	Lys
30	Leu	Val	Asp	Pro 20	Ile	Leu	Arg	Arg	His 25		Leu	Leu	Pro	Ser 30		Leu
	Lys	Arg	Ile 35	Ala	Val	Gly	Met	Phe 40	Phe	Val	Met	Cys	Ser 45	Ala	Phe	Ala
35	Ala	Gly 50	Ile	Leu	Glu	Ser	Lys 55	Arg	Leu	Asn	Leu	Val 60	Lys	Glu	Lys	Thr
40	Ile 65	Asn	Gln	Thr	Ile	Gly 70	Asn	Val	Val	Tyr	His 75	Ala	Ala	Asp	Leu	Ser 80
			Trp		85					90					95	
45			Ser	100					105					110		
50			Gln 115					120					125			
50		130	Phe				135					140				
55	145		Trp			150					155					160
			Asn		165					170					175	
60	Leu	Leu	Leu	Phe 180	Leu	Ile	Ile	Ser	Val 185	Lys	Tyr	Asp	His	His 190	Arg	Asp

	His	Gln	Arg 195	Ser	Arg	Ala .	Asn	Gly 7	Val :	Pro '	Thr		Arg 2 205	Arg A	Ala	
5																
	(2)	INFO	ORMAT	NOI	FOR	SEQ	ID N	10: 2	06:							
10				() () ()	A) LI B) T D) T	ENGTH YPE : OPOLO	H: 1 ami XGY:	ERIST 96 am no ac line PTION	mino cid ear	ació		206	5 :			
15	Met 1	Arg	Ser	Arg	Ile 5	Arg	Glu	Phe	Asp	Ser 10	Ser	Thr	Leu	Asn	Glu 15	Ser
20	Val	Arg	Asn	Thr 20	Ile	Met	Arg	Asp	Leu 25	Lys	Ala	Val	Gly	Lys 30	Lys	Phe
20	Met	His	Val 35	Leu	Tyr	Pro	Arg	Lys 40	Ser	Asņ	Thr	Leu	Leu 45	Arg	Asp	Trp
25	Asp	Leu 50		Gly	Pro	Leu	Ile 55	Leu	Cys	Val	Thr	Leu 60	Ala	Leu	Met	Leu
	Gln 65	Arg	, Asp	Ser	Ala	Asp 70	Ser	Glu	Lys	Asp	Gly 75	Gly	Pro	Gln	Phe	Ala 80
30	Glu	Val	Phe	· Val	Ile 85		Trp	Phe	Gly	Ala 90	Val	Thr	Ile	Thr	Leu 95	Asn
35	Ser	Lys	s Leu	Leu 100		Gly	Asn	Ile	Ser 105	Phe	Phe	Gln	Ser	Leu 110	Cys	Val
33	Leu	Gly	7 Tyr 115		Ile	. Leu	Pro	Leu 120		Val	Ala	Met	Leu 125	Ile	Cys	Arg
40	Leu	Va.		ı Leu	ı Ala	Asp	Pro 135	Gly	Pro	Val	Asn	Phe 140		Val	Arg	Leu
	Phe 145		l Val	l Il∈	e Val	150		e Ala	Trp	Ser	11e		Ala	Ser	Thr	Ala 160
45	Phe	e Le	u Ala	a Asp	Ser 169		n Pro) Pro) Asn	170		Ala	a Leu	. Ala	Val 175	Туг
50	Pro	o Va	1 Phe	e Leu 180		э Туг	: Pho	e Val	l Il∈ 185		Trp	Met	: Ile	190		Phe
50	Thi	r Pr	o Gl:		3											
55	(2	AI (IFORM	ATIO	N FO	R SE	Q ID	NO:	207	:						
			(i)	SEQ				TERI 331			ids					
60								nino								

(D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 207:

5	Met 1	Ala	Lys	Asp	Gln 5	Ala	Val	Glu	Asn	Ile 10	Leu	Val	Ser	Pro	Val 15	Val
	Val	Ala	Ser	Ser 20	Leu	Gly	Leu	Val	Ser 25	Leu	Gly	Gly	Lys	Ala 30	Thr	Thr
10	Ala	Ser	Gln 35	Ala	Lys	Ala	Val	Leu 40	Ser	Ala	Glu	Gln	Leu 45	Arg	Asp	Glu
15	Glu	Val 50	His	Ala	Gly	Leu	Gly 55	Glu	Leu	Leu	Arg	Ser 60	Leu	Ser	Asn	Ser
	Thr 65	Ala	Arg	Asn	Val	Thr 70	Trp	Lys	Leu	Gly	Ser 75	Arg	Leu	Tyr	Gly	Pro 80
20	Ser	Ser	Val	Ser	Phe 85	Ala	Asp	Asp	Phe	Val 90	Arg	Ser	Ser	Lys	Gln 95	His
	Tyr	Asn	Cys	Glu 100	His	Ser	Lys	Ile	Asn 105	Phe	Arg	Asp	Lys	Arg 110	Ser	Ala
25	Leu	Gln	Ser 115	Ile	Asn	Glu	Trp	Ala 120	Ala	Gln	Thr	Thr	Asp 125	Gly	Lys	Leu
30		130			Lys		135					140				
	145				Phe	150					155					160
35					Arg 165					170					175	
40				180	His				185					190		
40			195		Gln			200					205			
45		210					215					220				Leu
	225				Thr	230					235					240
50					Val 245					250					255	
				260	Gln				265					270		
55			275		Lys			280					285			
60	Leu	Туг 290	Leu	Ala	Ser	Val	Phe 295	His	Ala	Thr	Ala	Phe 300	Glu	Leu	Asp	Thr

	305	Gly	Asn	Pro) L∈		nr A 10	arg 1	.1e :	riir (319 C	31y \	/ai A	ug 1	3	320
5	Val :	Phe	туr	Ala	a As 32		is F	ro E	Phe :		Ser 3 330	(aa					
10	(2)				UEN((A) (B)	CE C LEI	CHARA NGTH PE:	ACTE : 58 amin	RIST ami o ac	ICS: .no a	ıcids						
15			(xi)	SE					line TION		Q ID	NO:	208	:			
	Met 1	Cys	Met	: Gl:	n L	eu E 5	Phe (Gly	Phe	Leu	Ala 10	Phe	Met	Ile 1	Phe I	Met 15	Cys
20	Trp	Val	Gly		V q 0	al 7	Tyr :	Pro	Val	Тут 25	Gln	Pro	Val	Gly :	Pro:	Lys	Gln
25	Tyr	Pro	Тут 35		n A	sn 1	Leu '	Tyr	Leu 40	Glu	Arg	Gly	Gly	Asp 45	Pro	Ser	Lys
25	Glu	Pro 50		ı Ar	g V	al '	Val	His 55	Tyr	Glu	Ile						
30	(2)	INF	ORM	ATIC	ON F	FOR	SEQ	ID 1	10: 2	209:							
35					(A (B (D) II) IV	ENGTI (PE : OPOL	H: 3 ami OGY:	92 a no a lin	cid ear	: aci EQ I		: 20	9 :			
40	Met 1		o Al	a Le	eu 7	Val 5	Glu	Asp	Asp	Ile	Cys 10	Ile	Leu	Asn	His	Glu 15	
	Ala	Hi:	s Ly		rg 2	Asp	Thr	Val	Thr	Pro 25		Ser	Ile	Tyr	Ser 30	Gly	Asp
45	Glu	ı Se		al A 85	la	Ser	His	Phe	Ala 40		ı Val	Thr	Ala	. Tyr 45	Glu	Asp	Ile
50	Lys		s Ar O	rg L	eu	Lys	Asp	Ser 55		ı Lys	s Glu	Asn	Ser 60	Leu	Leu	Lys	Lys
50	Arg		e Ai	rg P	he	Leu	Glu 70		Lys	Le	ı Ile	Ala 75		, Phe	Glu	Glu	Glu 80
55	Th	r Se	er Se	er V	al	Gly 85		Glu	ı Glr	n Va	l Asr 90		s Alá	a Tyr	His	Ala 99	
	Ar	g Gl	u V		:уs 100	Ile	Asp	Arg	j Asi	2 As:		ı Ly:	s Sei	r Lys	110		Lys
60	Me	t As	sn L	ys A	Asp	Asn	Ser	Glu	ı Se:	r Le	u Ly:	s Va	l Le	u Asr	ı Glu	ı Glı	n Leu

			115					120					125			
5	Gln	Ser 130	Lys	Glu	Val	Glu	Leu 135	Leu	Gln	Leu	Arg	Thr 140	Glu	Val	Glu	Thr
	Gln 145	Gln	Val	Met	Arg	Asn 150	Leu	Asn	Pro	Pro	Ser 155	Ser	Asn	Trp	Glu	Val 160
10	Glu	Lys	Leu	Ser	Cys 165	Asp	Leu	Lys	Ile	His 170	Gly	Leu	Glu	Gln	Glu 175	Leu
	Glu	Leu	Met	Arg 180	Lys	Glu	Cys	Ser	Asp 185	Leu	Lys	Ile	Glu	Leu 190	Gln	Lys
15	Ala	Lys	Gln 195	Thr	Asp	Pro	Tyr	Gln 200	Glu	Asp	Asn	Leu	Lys 205	Ser	Arg	Asp
20	Leu	Gln 210	Lys	Leu	Ser	Ile	Ser 215	Ser	Asp	Asn	Met	Gln 220	His	Ala	Tyr	Trp
	225					230					235			Gln		240
25					245					250				Lys	255	
				260					265					Ser 270		
30			275					280					285	Pro		
35		290					295					300		Pro		
	305					310					315			Ser		320
10					325					330				Gln	335	
15				340					345					Val 350		
+3			355					360					365	Lys		
50		370					375		Pro	Leu	His	Tyr 380	Leu	Asp	Gln	His
	385	Gln	Asn	Cys	Leu	Тут 390	Lys	Asn								
55	(2)	INFC	RMAT	NOI	FOR	SEQ	ID N	IO: 2	10:							
		•	(i) S					ERIST								
50								7 am: no ac		acid	5					

```
(D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 210:
     Met His His His Thr Gln Leu Met Phe Ile Tyr Leu Phe Ile Tyr Leu
5
                                          10
     Phe Ile Leu Gly Val Phe Phe Phe Phe Xaa
                  20
10
      (2) INFORMATION FOR SEQ ID NO: 211:
             (i) SEQUENCE CHARACTERISTICS:
15
                    (A) LENGTH: 39 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 211:
      Met Asn Cys Ile Leu Leu Leu Tyr Leu Leu Ile Pro Thr Ile Ser Ile
20
                                          10
      Ser Val Val Pro Tyr Val Ala Leu Asn Ile Lys Tyr Ile Lys Glu Cys
                  2.0
25
      Thr Glu Asn Ser Phe Tyr Xaa
30
      (2) INFORMATION FOR SEQ ID NO: 212:
              (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 71 amino acids
35
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 212:
      Met Leu Leu His Leu Thr Ala Ala Phe Leu Gln Arg Ala Gln Phe Ser
40
      Thr Tyr Phe Pro Gly Tyr Phe Asp Gly Gln Tyr Trp Leu Trp Trp Val
                                        25
                                                            30
       Phe Leu Val Leu Gly Phe Leu Leu Phe Leu Arg Gly Phe Ile Asn Tyr
45
       Ala Lys Val Arg Lys Met Pro Glu Thr Phe Ser Asn Leu Pro Arg Thr
            50
                              55
 50
       Arg Val Leu Phe Ile Tyr Xaa
                            70
        65
 55
       (2) INFORMATION FOR SEQ ID NO: 213:
               (i) SEQUENCE CHARACTERISTICS:
                      (A) LENGTH: 83 amino acids
                      (B) TYPE: amino acid
 60
```

```
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 213:
      Met Leu Thr Phe Phe Met Ala Phe Leu Phe Asn Trp Ile Gly Phe Phe
  5
      Leu Ser Phe Cys Leu Thr Thr Ser Ala Ala Gly Arg Tyr Gly Ala Ile
                                       25
10
      Ser Gly Phe Gly Leu Ser Leu Ile Lys Trp Ile Leu Ile Val Arg Phe
      Ser Thr Tyr Phe Pro Ala Phe Met Asn Ser Leu Ser Arg Ser Lys Arg
15
      Thr Pro Ala Gly Ser Glu Ser Arg Cys Arg Thr Gln Arg Asn Asn His
                          70
      Leu Leu Xaa
20
      (2) INFORMATION FOR SEQ ID NO: 214:
25
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 81 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
30
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 214:
      Met Ser Lys Arg Ser Ala Ser Phe Ile Leu Leu Pro Leu Leu Phe Leu
                                          10
35
      Lys Gly Ser Phe Ala Lys Leu Asn Ala Arg Ile Ser Asp Cys Leu Glu
      Glu Arg Tyr Cys His Asn Leu Trp Met Val Phe Gln Gly Cys Val Ile
40
      Thr Glu Leu His Leu Ser Arg Met Ser Lys Thr Leu Ser Ser Leu Cys
      Tyr Asp Phe Val Ile Asn Val Tyr Ile Phe Phe Lys Phe Leu Asp Ile
45
                           70
      Thr
50
      (2) INFORMATION FOR SEQ ID NO: 215:
             (i) SEQUENCE CHARACTERISTICS:
55
                    (A) LENGTH: 49 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 215:
60
      Met Cys Ser Leu Phe Glu Ser Arg Phe Phe Cys Phe Val Leu Phe Ser
```

(D) TOPOLOGY: linear

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	1				5					10					15	
	Glu L	ys I	Ile I	[le 0	3ln I	Leu (Cys A	Ala	Ser 25	Ile A	Ala :	Phe I	Leu	Cys :	Phe '	Val
5	Lys H	is V	Val 1 35	Pro 1	rp l	Pro 1	Lys '	Frp	Lys .	Arg :	Lys (Cys 1	Leu 45	Ile .	Asn .	Ala
10	Phe															
15	(2) I			EQUE (A	NCE	CHAR INGTH	ACTE	RIST	TICS:		ls					
20		((xi)	(E (E SEQU) TC	PE: POLC DES	GY:	line		EQ II) NO:	: 216	5:			
	Met 7	Thr	Leu	Arg	Pro 5	Ser	Leu	Leu	Pro	Leu 10	His	Leu	Leu	Leu	Leu 15	Leu
25	Leu l	Leu	Ser	Ala 20	Ala	Val	Cys	Arg	Ala 25	Glu	Ala	Gly	Leu	Glu 30	Thr	Glu
20	Ser	Pro	Val 35	Arg	Thr	Leu	Gln	Val 40	Glu	Thr	Leu	Val	Glu 45	Pro	Pro	Glu
30	Pro	Cys 50	Ala	Glu	Pro	Ala	Ala 55	Phe	Gly	Asp	Thr	Leu 60	His	Ile	His	Tyr
35	Thr 65	Gly	Ser	Leu	Val	Asp 70	Gly	Arg	Ile	Ile	Asp 75	Thr	Ser	Leu	Thr	Arg 80
	Asp	Pro	Leu	Val	Ile 85	Glu	Leu	Gly	Gln	Lys 90	Gln	Val	Ile	Pro	Gly 95	Leu
40	Glu	Gln	Ser	Leu 100	Leu	Asp	Met	Cys	Val 105	Gly	Glu	Lys	Arg	Arg 110		Ile
45	Ile	Pro	Ser 115	His	Leu	Ala	Tyr	Gly 120		Arg	Gly	Phe	Pro 125		Ser	Val
4 5	Pro	Ala 130		Ala	Val	Val	Gln 135		Asp	Val	Glu	Leu 140		e Ala	. Leu	Ile
50	Arg 145	Ala	. Asn	Tyr	Trp	Leu 150		: Leu	ı Val	. Lys	Gly 155		. Leu	ı Pro	Leu	160
	Gly	Met	. Ala	Met	Val 165		Pro	Sei	r Trp	170		Leu	Gly	/ Il∈	e Thr 175	Tyr
55	Thr	Glu	ı Arg	180		e Asp	Pro	Ly:	5 Sei 189		Lys	s Arg	g Se:	r Sei 190		g Lys
60	Arg	Asr	195		Arç	g Alá	a Lys	200	g Ası O	n Ası	ı Ly:	5				

	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	217:							
5			(i)	((A) I (B) T		TH: 1 ami	.86 a .no a			ds					
10	Met	Lvs		SEQ	UENC	E DE	SCRI	PTIO	N: S Pro					_	2	_,
	1				5					10					15	
15	Ser	Ile	Ser	Leu 20	Trp	Ile	Ile	Ala	Ala 25	Trp	Thr	Val	Arg	Val 30	Cys	Glu
	Ser	Pro	Glu 35	Ser	Pro	Ala	Gln	Pro 40	Ser	Gly	Ser	Ser	Leu 45	Pro	Ala	Trp
20	Tyr	His 50	ąsA	Gln	Gln	Asp	Val 55	Thr	Ser	Asn	Phe	Leu 60	Gly	Ala	Met	Trp
25	Leu 65	Ile	Ser	Ile	Thr	Phe 70	Leu	Ser	Ile	Gly	Tyr 75	Gly	Asp	Met	Val	Pro 80
	His	Thr	Tyr	Cys	Gly 85	Lys	Gly	Val	Cys	Leu 90	Leu	Thr	Gly	Ile	Met 95	Gly
30	Ala	Gly	Cys	Thr 100	Ala	Leu	Val	Val	Ala 105	Val	Val	Ala	Arg	Lys 110	Leu	Glu
	Leu	Thr	Lys 115	Ala	Glu	Lys	His	Val 120	His	Xaa	Phe	Met	Met 125	Asp	Thr	Gln
35	Leu	Thr 130	Lys	Arg	Ile	Lys	Asn 135	Xaa	Ala	Ala	Asn	Val 140	Leu	Xaa	Glu	Thr
40	Trp 145	Leu	Ile	Tyr	Lys	His 150	Thr	Lys	Leu	Leu	Lys 155	Lys	Ile	Asp	His	Ala 160
	Lys	Val	Arg	Asn	Thr 165	Arg	Gly	Ser	Ser	Ser 170	Lys	Tyr	Pro	Pro	Val 175	Glu
45	Glu	Arg	Gln	Asp 180	Gly	Thr	Glu	Glu	Ala 185	Glu						
50	(2)	INFO	ORMAT	NOI	FOR	SEQ	ID N	IO: 2	18:							
			(i) \$	(.	A) L		H: 9	am:	TICS: ino a cid		5					
55			(xi)			OPOLO E DES			ear J: SE	EQ IE	ONO:	: 218	3:			
	Met 1	Lys	Phe	Leu	Ala 5	Val	Leu	Val	Leu	Leu 10	Gly	Val	Ser	Ile	Phe 15	Leu
60	Val	Ser	Ala	Gln	Asn	Pro	Thr	Thr	Ala	Ala	Pro	Ala	Asp	Thr	тут	Pro

				20					25					30		
	Ala	Thr	Gly 35	Pro	Ala	Asp	Asp	Glu 40	Ala	Pro	Asp	Ala	Glu 45	Thr	Thr	Ala
5	Ala	Ala 50	Thr	Thr	Ala	Thr	Thr 55	Ala	Ala	Pro	Thr	Thr 60	Ala	Thr	Thr	Ala
10	Ala 65	Ser	Thr	Thr	Ala	Arg 70	Lys	Asp	Ile	Pro	Val 75	Leu	Pro	Lys	Trp	Val 80
	Gly	Asp	Leu	Pro	Asn 85	Gly	Arg	Val	Cys	Pro 90						
15																
	(2)	INF	ORMA'													
20				- ((A) L B) T D) T	ENGT YPE : OPOL	'H: 1 ami OGY:	ERIS 39 a no a lin PTIO	mino cid ear	aci		: 21	9:			
25	Met 1		Ser	Ala	Ala 5	Ala	Asp	His	Trp	Ala 10	Trp	Leu	Leu	Val	Leu 15	Ser
	Phe	· Val	. Phe	Gly 20		Asn	Val	Leu	Arg 25	Ile	Leu	Leu	Pro	Ser 30	Phe	Ser
30	Ser	Phe	Met 35		Arg	Val	Leu	Gln 40		Asp	Ala	Glu	Gln 45	Glu	Ser	Gln
35	Met	Arg 50	g Ala)	Glu	Ile	Gln	Asp 55		Lys	Gln	Glu	Leu 60		Thr	Val	Asr
	Met 65		asp	Glu	Phe	Ala 70		Tyr	· Ala	Arg	Leu 75		Arg	Lys	Ile	Asr 80
40	Lys	s Mei	t Thr	: Asr	Lys 85		ı Lys	Thr	His	90		Ala	Arg	Thr	Ala 95	
45	Let	ı Ala	a Lys	100		Tr	Va.	l Ile	Ser 105		L Ala	a Ph∈	э Туг	Val		ı Glr
45	Ala	a Al	a Lev 115		: Ile	e Sei	r Lei	ı Ile 120) Lys	з Тул	туз	Ser 125		. Pro	o Vai
50	Ala	a Va 13	1 Va:	l Pro	o Sei	c Ly:	s Try		e Thi	c Le	ı Xaa	a				
55	(2) IN	FORM	ATIO	N FO	R SE	Q ID	NO:	220	:	٠					
_			(i)	SEQ	(A) (B)	LENC TYPE	TH: E: an	TERI 48 a	mino acid	aci l	.ds					
60			1200	۱ د	OTTEN			?: li			TD N	iO · 2	20 -			

	Met 1	Ser	Ser	Ala	Ala 5	Ala	Asp	His	Trp	Ala 10	Trp	Leu	Leu	Val	Leu 15	Ser
5	Phe	Val	Phe	Gly 20	Cys	Asn	Val	Leu	Arg 25	Ile	Leu	Leu	Pro	Ser 30	Phe	Ser
10	Ser	Phe	Met 35	Ser	Arg	Val	Leu	Gln 40	Lys	Asp	Ala	Asp	Arg 45	Ser	His	Arg
15	(2)	INF	ORMA'	TION	FOR	SEQ	ID I	NO: 2	221:							
20				(A) L B) T D) T	ENGT YPE: OPOL	H: 7 ami OGY:	0 am no a lin	ino cid ear	acid		: 22	1:			
25	Met 1	Thr	Ala	Pro	Leu 5	Pro	Pro	Leu	Ser	Gly 10	Leu	Ala	Leu	Phe	Leu 15	Ile
	Val	Phe	Phe	Ser 20	Leu	Gly	Val	Phe	Cys 25	Ile	Cys	His	Ser	His 30	Trp	Tyr
30	His	Thr	Leu 35	Gln	Gln	Met	Ala	Gly 40	Thr	Glu	Pro	Lys	Ala 45	Leu	Leu	Leu
35	Ser	Pro 50	Pro	Ala	Ala	Thr	Thr 55	Phe	Val	Thr	Val	Thr 60	His	Glu	Val	Trp
	Lys 65	Glu	Gln	Ala	Leu	Ala 70										
40	(2)			rion												
45				(A) L B) T D) T	ENGT: YPE : OPOL	H: 8 ami: OGY:	3 am no a lin	ino a cid ear	acid		: 222	2:			
50	Met 1	Thr	Cys	Ser	Val 5	Ala	Leu	Leu	Leu	Ile 10	Leu	Gly	Leu	Arg	Cys 15	Ser
	Gly	Val	Arg	Pro 20	Gly	Leu	Val	Gly	Glu 25	Gly	His	Asn	Pro	Ser 30	Leu	Leu
55	Val	Cys	Leu 35	Leu	Leu	Lys	Asp	Ser 40	Arg	Thr	Asn	Gln	Gly 45	Ser	Cys	Pro
60	Gly	Gly 50	Pro	Trp	Ser	Glu	Arg 55	Asp	Ile	Glu	Ser	Val 60	Thr	Ser	Asp	Asn

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	Cys (Glu	Ala	Thr	Leu	Gly 70		r A	rg A	sn H		er L	eu P	ro S	er A	sn T	80 80
5	Tyr A	Asn	Ser														
10	(2)			SEQU	ENCE	E CHI	ARA(TH:	CTEF 43	: 22 RISTI amir	CS:	cids						
15			(xi)		(D) '	TOPO	LOG	Y: :	linea	ar	Q ID	NO:	223	:			
	Met 1	Leu	Thr	Arg		c Le	a L	ys 1	hr I	eu I	Pro S 10	Ser A	ala (Cys :	Thr A	Ala : 15	Phe
20	Leu	Leu	Leu	Phe 20		e Le	u P	he S	Ser S	Ser (25	Gly A	Asp I	?ro (Glu 1	Leu : 30	Ser	Cys
25	Ser	Cys	Thr		ı Ar	g Th	r G	ln s	Ser S 40	Ser '	Trp :	Ser					
30	(2)	INF	(i)	SEQ	UENC (A) (B) (D)	E CH LENG TYPI TOPG	IARI STH E: S	ACTE : 18 amir GY:	no ac line	ICS: ino id ar	acio		224				
35	Met 1										EQ II Leu 10				Arg	His 15	Gly
40	Ala	Gli	n Gl		s Pr	co Se	er 1	Pro	Asp	Ala 25	Gly	Pro	His	Gly	Gln 30	Gly	Arg
	Val	Hi		n Al 5	a A.	la P:	ro:	Leu	Ser 40	Asp	Ala	Pro	His	Asp 45	Asp	Ala	His
45	Gly	As:		ie Gl	n T	yr A	sp	His 55	Glu	Ala	Phe	Leu	Gly 60	Arg	Glu	Val	Ala
50	Lys 65		u Ph	ne As	sp G		eu 70	Thr	Pro	Glu	Glu	Ser 75	Gln	Ala	Arg	Leu	Gly 80
30	Arg	g Il	e Va	al A		rg M 85	let	Asp	Arg	Ala	Gly 90		Gly	Asp	Gly	Trp 95	Val
55	Se:	r Le	eu A		lu L 00	eu A	rg	Ala	Trp	Ile 105		His	Thr	Gln	Gln 110		, His
	Il	e Ar		sp S 15	er V	al S	Ser	Ala	Ala 120		asp	Thr	Tyr	Asp 125		: Asp	Arg
60	As	ກ G:	lv A	ra V	al G	aly :	rp	Glu	ı Glu	Leu	ı Arç	Asr	. Xaa	t Thi	туг	Gly	y His

		130					135					140				
5	Xaa 145		Pro	Xaa	Glu	Glu 150	Phe	His	Asp	Val	Glu 155	Asp	Ala	Glu	Thr	Tyr 160
3	Lys	Lys	Met	Leu	Xaa 165	Arg	Asp	Glu	Arg	Arg 170	Phe	Arg	Val	Ala	Asp 175	Gln
10	Asp	Gly	Asp	Ser 180	Met	Ala	Thr	Arg								
15	(2)	INF	ORMA	SEQU	ENCE	CHAI	RACT:	ERIS'	rics							
20			(xi)	(B) T D) T	ENGT YPE: OPOL E DE:	ami OGY:	no a lin	cid ear			: 22	5:			
	Met 1	Trp	Leu	Phe	Ile 5	Leu	Leu	Ser	Leu	Ala 10	Leu	Ile	Ser	Asp	Ala 15	Met
25	Val	Met	Asp	Glu 20	Lys	Val	Lys	Arg	Ser 25	Leu	Cys	Trp	Thr	Arg 30	Leu	Leu
30	Pro	Ser	Ala 35	Thr	Thr	Met	Pro	Хаа 40	Thr	Arg	Ile	Thr	Pro 45	Asn	Thr	Gly
	Ala	Glu 50	Xaa	Ile	Ser	Val	Xaa 55	Thr	Ala	Thr	Ser	Ser 60	Pro	Ser	Pro	Leu
35	Thr 65	Ala	Pro	Ile	Met	Trp 70	Pro									
40	(2)		ORMAI													
			(i) :	() ()	A) L B) T	ENGTI YPE : OPOLA	H: 1 ami	0 am no a	ino a		5					
45			(xi)							EQ II	ONO:	: 226	5 :			
	Met 1	His	Val	Phe	Val 5	Leu	Glu	Ile	Phe	Leu 10						
50																
	(2)		ORMAT													
55			(i)	() () ()	A) L: B) T D) T	ENGTI YPE : OPOLA	H: 1 amin OGY:	38 ar no ao line	mino cid ear	acio		: 225	7 :			
50	Met		Val							_				I 011	A1-	T

	1				5					10					15		
	Thr	Phe	Ile	Thr 20	Asp	Asn	Ser	Leu	Val 25	Ala	Ala	Gly	His	Asp 30	Cys	Ph	ie
5	Pro	Val	Leu 35	Phe	Thr	Tyr	Asp	Ala 40	Ala	Ala	Gly	Met	Leu 45	Ser	Phe	G1	·Y
10	Gly	Arg 50	Leu	Asp	Val	Pro	Lys 55	Gln	Ser	Ser	Gln	Arg 60	Gly	Leu	Thr	Al	.a
	Arg 65	Glu	Arg	Phe	Gln	Asn 70	Leu	Asp	Lys	Lys	Ala 75	Ser	Ser	Glu	Gly		Ly 30
15	Thr	Ala	Ala	Gly	Ala 85	Gly	Leu	Asp	Ser	Leu 90	His	Lys	Asn	Ser	Val 95		er
20	Gln	Ile	Ser	Val 100	Leu	Ser	Gly	Gly	Lys 105		Lys	Суѕ	Ser	Gln 110	Phe	e C <u>y</u>	ys
20	Thr	Thr	Gly 115		Asp	Gly	Gly	Met 120	Ser	Ile	Trp	Asp	Val 125		Ser	: L	eu
25	Glu	Ser 130	Ala	Leu	Lys	Asp	Leu 135		Ile	Lys							
30	(2)	INF	ORMA	SEQU	FOR JENCE (A) ! (B) '	E CHA	ARACT	TERIS 23 au	STICS mino	S: aci	ds						
35			(xi)		(D) '					SEQ :	ID N	0: 2:	28:				
		ı Gly l	y Sei	c Let		Thr	Alá	a Pro	Sei	Sei		a Lei	ı Pro	Th:	c Le		ly
40	Ala	a Arg	g Arq	g Thi		g Sei	. Lys	5									
45	(2) IN	FORM	OITA	N FO	R SE	Q ID	NO:	229	:							
50					(B)	LENC TYPE TOPO	TH: : an LOGY	133 nino : li	amir acid near	no ac 1 :		10: 2	229:				
55	Me	t Th	ır Ty	r Ph	e Se	r Gl 5	y Le	u Le	u Va		.e L∈ .0	eu Al	a Ph	ne Al		la ' l5	Trp
	Va	l Al	.a L∈		.a Gl !0	u Gl	y Le	eu Gl		al Al 25	a Va	al Ty	r Al		.a A:	la	Val
60	L€	eu Le	eu Gl	ly A] 35	la Gl	.у Су	's Al		r Il 10	le Le	eu Va	al Th		er Le 15	eu A	la	Met

	Thr	Ala 50	Asp	Leu	Ile	Gly	Pro 55	His	Thr	Asn	Ser	Gly 60	Ala	Phe	Val	Tyr
5	Gly 65	Ser	Met	Ser	Phe	Leu 70	Asp	Lys	Val	Ala	Asn 75	Gly	Leu	Ala	Val	Met 80
10	Ala	Ile	Gln	Ser	Leu 85	His	Pro	Cys	Pro	Ser 90	Glu	Leu	Cys	Cys	Arg 95	Ala
	Cys	Val	Ser	Phe 100	Tyr	His	Trp	Ala	Met 105	Val	Ala	Val	Thr	Gly 110	Gly	Val
15	Gly	Val	Ala 115	Ala	Ala	Leu	Cys	Leu 120	Суѕ	Ser	Leu	Leu	Leu 125	Trp	Pro	Thr
	Arg	Leu 130	Arg	Arg	Xaa											
20	(0)															
	(2)			rion												
25				(1 (1	A) L: B) T D) T	ENGT YPE : OPOLA	H: 2: ami: OGY:	8 am no a lin	ino d cid ear	: acid: EQ II		: 23():			
30	Gly 1	Lys	Pro	Thr	Gly 5	Lys	Ser	Leu	Pro	Leu 10	Met	Trp	Met	Ile	Leu 15	Met
35	Gln	Pro	Ile	Ile 20	Met	Ile	Ser	Met	Met 25	Ser	Asn	Gly				
40	(2)	INFO	RMAT	NOI	FOR	SEQ	ID N	10: 2	:31:							
40		,	(i) S	(E	A) LI 3) T	ENGTI PE:	1: 61 amir	lam:	ino a cid	: acids	3					
45		((xi)			OPOLO E DES				EQ IE	NO:	231	:			
	Met 1	Gln	Gly	Lys	Phe 5	Met	Lys	Val	Gln	Val 10	Tyr	Arg	Phe	Leu	Lys 15	Tyr
50	Leu	Leu	Met	Leu 20	Leu	Cys	Met	Phe	Val 25	Asn .	Arg	Gly	Met	Ser 30	Lys	Asp
	Ser	Thr	Lys 35	Lys	Pro	Gly	Gln	Glu 40	Lys	Leu	Lys	Val	Ser 45	Leu	Gly	Ser
55	Ile	Leu 50	Asn	Met	Lys	Ser	Gln . 55	Arg	Pro	Leu	Ser	Trp (Cys			
50	(2)	INFO	RMAT	ION :	FOR	SEQ	ID N	0: 2	32:							

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(i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 29 amino acids
                    (B) TYPE: amino acid
5
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 232:
     Met Met Glu Arg Ser Met Met Ile Leu Leu Met Ala Ala Ser Met Thr
10
     Met Thr Ser Thr Gln Leu Trp Ser Phe Cys Cys Val His
                                      25
                   20
15
      (2) INFORMATION FOR SEQ ID NO: 233:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 18 amino acids
                     (B) TYPE: amino acid
20
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 233:
      Met Trp Tyr Gln Leu Ala Lys Glu Glu Pro Gly Val Gly Ala Cys Ala
25
                       5
                                           10
      Leu Asp
30
       (2) INFORMATION FOR SEQ ID NO: 234:
              (i) SEQUENCE CHARACTERISTICS:
35
                     (A) LENGTH: 2 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 234:
40
       Leu Xaa
        1
       (2) INFORMATION FOR SEQ ID NO: 235:
 45
              (i) SEQUENCE CHARACTERISTICS:
                      (A) LENGTH: 72 amino acids
                      (B) TYPE: amino acid
 50
                      (D) TOPOLOGY: linear
               (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 235:
       Met Leu Ile Cys Arg Leu Val Leu Leu Ala Asp Pro Gly Pro Val Asn
                                             10
 55
       Phe Met Val Arg Leu Phe Val Val Ile Val Met Phe Ala Trp Ser Ile
       Val Ala Ser Thr Ala Phe Leu Ala Asp Ser Gln Pro Pro Asn Arg Arg
 60
                                     40
```

	Ala	Leu 50	Ala	Val	Tyr	Pro	Val 55	Phe	Leu	Phe	Tyr	Phe 60	Val	Ile	Ser	Trp
5	Met 65		Leu	Thr	Phe	Thr 70	Pro	Gln								
10	(2)	INF	ORMA	rion	FOR	SEQ	ID I	NO: :	236:							
15			(i) (xi)	(A) L B) T D) T	ENGT YPE: OPOL	H: 9 ami OGY:	6 am no a lin	ino cid ear	acid		: 23	6 :			
20	Met 1	Arg	Ser	Leu	Leu 5	Leu	Leu	Ser	Ala	Phe 10	Cys	Leu	Leu	Glu	Ala 15	Ala
	Leu	Ala	Ala	Glu 20	Val	Lys	Lys	Pro	Ala 25	Ala	Ala	Ala	Ala	Pro 30	Gly	Thr
25	Ala	Glu	Lys 35	Leu	Ser	Pro	Lys	Ala 40	Ala	Thr	Leu	Ala	Glu 45	Arg	Xaa	Pro
	Ala	Trp 50	Pro	Ser	Ala	Cys	Thr 55	Arg	Pro	Trp	Pro	Arg 60	Thr	Arg	Gln	Trp
30	Arg 65	Thr	Ser	Trp	Cys	His 70	Pro	Trp	Trp	Trp	Pro 75	Arg	Arg	Trp	Gly	Ser 80
35	Cys	Arg	Trp	Ala	Ala 85	Arg	Arg	Pro	Arg	Arg 90	Arg	Arg	Pro	Arg	Gln 95	Cys
40	(2)		ORMAT													
45			(i) 5 (xi)	(. (: (:	A) Li B) T D) T	ENGTI YPE: OPOLA	H: 14 amin OGY:	43 ar no ao line	mino cid ear	aci		: 237	7:			
50	Met 1	Arg	Ser	Leu	Leu 5	Leu	Leu	Ser	Ala	Phe 10	Cys	Leu	Leu	Glu	Ala 15	Ala
	Leu	Ala	Ala	Glu 20	Val	Lys	Lys	Pro	Ala 25	Ala	Ala	Ala	Ala	Pro 30	Gly	Thr
55	Ala	Glu	Lys 35	Leu	Ser	Pro	Lys	Ala 40	Ala	Thr	Leu	Ala	Glu 45	Arg	Lys	Arg
60	Pro	Gly 50	Leu	Gln	Leu	Val	Pro 55	Gly	His	Gly	Gln	Gly 60	Pro	Gly	Ser	Gly

	G1u 65	His	Pro	Gly	Val	Thr 70	Arg	Gly	Gly	Gly	Leu 75	Val	Ala	Gly	Ala	Arg 80
5	Val	Ala	Gly	Arg	Gln 85	Gly	Asp	His	Gly	Val 90	Ala	Gly	Gln	Gly	Ser 95	Ala
	Glu	Arg	Arg	Ala 100	Ala	Ala	Arg	Arg	Gly 105	Gly	Ala	Arg	Arg	Pro 110	Gly	Arg
10	Ala	Ala	Ala 115	Leu	Thr	Gln	Gln	Leu 120	His	Gly	Ala	Gln	Arg 125	Asp	Leu	Glu
15	Ala	Gly 130	Gln	Pro	Thr	Val	Arg 135	Thr	Gln	Leu	Ser	Glu 140	Leu	Arg	Xaa	
20	(2)	INF	(i)	(ENCE (A) L (B) T (D) T	CHA ENGT YPE:	RACT TH: 1 the aming and a contraction and a con	ERIS 142 a ino a : lir	TICS amino acid near	aci		v. 22	۵.			
25	Met 1	_	(xi) Ser			Leu			N: S Ala		. Cys			Glu	Ala 15	Ala
30	Leu	. Ala	a Ala	Glu 20		Lys	Lys	Pro	Ala 25		ı Ala	. Ala	Ala	Pro 30		Thr
	Ala	Glu	1 Lys 35		ser	Pro	Lys	40		Thr	: Leu	ı Ala	Glu 45		Xaa	Arg
35	Pro	Gl ₃		ı Glr	ı Lev	ı Val	l Pro		/ His	Gly	/ Glr	Gly 60		Gly	Ser	Gly
40	Glu 65		s Pro	o Gly	/ Val	. Thi		g Gly	y Gly	/ Gly	y Let 75		. Ala	Gly	Ala	Arg 80
10	Va]	L Ala	a Gl	y Arg	g Glr 85		y As	o His	s Gly	y Va.		a Gly	/ Glr	ı Gly	95	Ala
45	Glı	ı Ar	g Ar	g Ala 10		a Al	a Ar	g Ar	g Gly 109		y Ala	a Ar	g Arg	110		y Arg
	Ala	a Al	a Al 11		u Th	r Gl	n Gl	n Le		a Gl	y Al	a Gl	n Arg) Le	u Glu
50	Al	a Gl 13		n Pr	o Th	r Va	1 Ar 13		r Gl	n Le	u Se	r Gl		ı Arg	g	
55	(2) IN							239							
			(i)	SEÇ	(A)	LEN	GTH:	54 a	STIC amino acio	ac	ids					
60									inear							

			(XI)	SEQ	OEINC	e De	SCRI	PHO	N: S	EQ I	טא ס	: 23	9:			
5	Asp 1	Pro	Glu	Ala	Ala 5	Asp	Ser	Gly	Glu	Pro 10	Gln	Asn	Lys	Arg	Thr 15	Pro
,	Asp	Leu	Pro	Glu 20	Glu	Glu	Tyr	Val	Lys 25	Glu	Glu	Ile	Gln	Glu 30	Asn	Glu
10	Glu	Ala	Val 35	Lys	Lys	Met	Leu	Val 40	Glu	Ala	Thr	Arg	Glu 45	Phe	Glu	Glu
	Val	Val 50	Val	Asp	Glu	Ser										
15																
	(2)			rion												
20			(i)	(A) L B) T	CHAI ENGT YPE : OPOL	H: 6 ami	3 am no a	ino cid		s					
			(xi)	SEQ						EQ I	D N O	: 24	0 :			
25	Gln 1	Lys	Leu	Lys	Arg 5	Lys	Ala	Glu	Glu	Asp 10	Pro	Glu	Ala	Ala	Asp 15	Ser
30	Gly	Glu	Pro	Gln 20	Asn	Lys	Arg	Thr	Pro 25	Asp	Leu	Pro	Glu	Glu 30	Glu	Туг
	Val	Lys	Glu 35	Glu	Ile	Gln	Glu	Asn 40	Glu	Glu	Ala	Val	Lys 45	Lys	Met	Leu
35	Val	Glu 50	Ala	Thr	Arg	Glu	Phe 55	Glu	Glu	Val	Val	Val 60	Asp	Glu	Ser	
	(2)	INF	ORMA!	rion	FOR	SEQ	ID 1	1 0: 2	241:							
10				SEQUI	ENCE	CHAI	RACTI	ERIS	rics							
				C	B) T	ENGT YPE:	ami:	no a	cid	aci	ds					
15			(xi)	SEQ		OPOL				EQ II	ON C	: 24	1:			
	Lys 1	Ala	Met	Glu	Lys 5	Ser	Ser	Leu	Thr	Gln 10	His	Ser	Trp	Gln	Ser 15	Leu
50	Lys	Asp	Arg	Tyr 20	Leu	Lys	His	Leu	Arg 25	Gly	Gln	Glu	His	Lys 30	Tyr	Leu
55	Leu	Gly	Asp 35	Ala	Pro	Val	Ser	Pro 40	Ser	Ser	Gln	Lys	Leu 45	Lys	Arg	Lys
,,,	Ala	Glu 50	Glu	Asp	Pro	Glu	Ala 55	Ala	Asp	Ser	Gly	Glu 60	Pro	Gln	Asn	Lys
50	Arg 65	Thr	Pro	Asp	Leu	Pro 70	Glu	Glu	Glu	Tyr	Val 75	Lys	Glu	Glu	Ile	Glr.

	Glu i	Asn	Glu	Glu	Ala 85	Val	Lys	Lys	Met	Leu 90	Val	Glu	Ala	Thr .	Arg 95	Glu
5	Phe (Glu	Glu	Val 100	Val	Val	Asp	Glu	Ser 105	Pro	Pro	Asp	Phe	Glu 110	Ile	His
10	Ile															
	(2)	TNIE	ימשפר	TION	FOR	SEO	TD N	JO: 2	242 :							
15	(2)	1111	(i)	SEQUI ()	ENCE A) Li B) T	CHAI ENGTI YPE : OPOL	RACTI H: 1 ami OGY:	ERIS 48 a no a lin	rICS: mino cid ear	aci		: 24	2:			
20	Leu 1	Pro	Ser	Tyr	Asp 5	Glu	Ala	Glu	Arg	Thr 10	Lys	Ala	Glu	Ala	Thr 15	Ile
25	Pro	Leu	Val	Pro 20	Gly	Arg	Asp	Glu	Asp 25	Phe	Val	Gly	Arg	Asp 30	Asp	Phe
	Asp	Asp	Ala 35	Asp	Gln	Leu	Arg	Ile 40	Gly	Asn	Asp	Gly	11e 45	Phe	Met	Leu
30	Thr	Phe 50		e Met	Ala	Phe	Leu 55		Asn	Trp	Ile	Gly 60	Phe	Phe	Leu	Ser
35	65	_		ı Thr		70					75					80
				ı Ser	85					90					95	i
40				0 Gly 100)				105					110		
	Leu	Va.	1 Let	ս Glչ 5	/ Phe	e Leu	Lev	120		Ar <u>c</u>	, Gl	/ Phe	125		тут	. Ala
45	Lys	13		g Lys	s Met	Pro	135		. Phe	e Ser	Ası	140		Arg	Thi	Arg
50	Val 145		u Ph	e Ile	Э											
	(2)	IN	FORM	ATIO	N FOI	R SE	Q ID	NO:	243	:						
55			(i)	SEQ	(A) (B)	LENG TYPE	TH: : an	24 a nino	STIC: mino acid near	aci	.ds					
60			(xi	i) SE							ID N	10: 2	43:			

	Ala 1		/ Arg	Tyr	Gly 5		Ile	Ser	Gly	Phe 10		Leu	Ser	Leu	Ile 15	Lys
5	Trp) Ile	e Leu	Ile 20		Arg	Phe	Ser								
10	(2)	INF	ORMA	SEQU	ENCE	CHA	RACT	ERIS	TICS ino		ls					
15			(xi)	SEQ	D) I					EQ I	D NO	: 24	4:			
	Met 1	Lys	His	Leu	Ser 5	Ala	Trp	Asn	Phe	Thr 10	Lys	Leu	Thr	Phe	Leu 15	Gln
20	Leu	Trp	Glu	Ile 20	Phe	Glu	Gly	Ser	Val 25	Glu	Asn	Cys	Gln	Thr 30	Leu	Thr
25	Ser	Tyr	Ser 35	Lys	Leu	Gln	Ile	Lys 40	Tyr	Thr	Phe	Ser	Arg 45	Gly	Ser	Thr
	Phe	Tyr 50	Ile													
30	(2)		ORMA(
35			(xi)	(A) L B) T D) T	ENGT YPE : OPOL	H: 2 ami OGY:	13 a no a lin	mino cid ear	aci		: 24	5:			
40	Phe 1	Ser	Ser	Asp	Phe 5	Arg	Thr	Ser	Pro	Trp 10	Glu	Ser	Arg	Arg	Val 15	Glu
	Ser	Lys	Ala	Thr 20	Ser	Ala	Arg	Cys	Gly 25	Leu	Trp	Gly	Ser	Gly 30	Pro	Arg
45	Arg	Arg	Pro 35	Ala	Ser	Gly	Met	Phe 40	Arg	Gly	Leu	Ser	Ser 45	Trp	Leu	Gly
50	Leu	Gln 50	Gln	Pro	Val	Ala	Gly 55	Gly	Gly	Gln	Pro	Asn 60	Gly	Asp	Ala	Pro
	Pro 65	Glu	Gln	Pro	Ser	Glu 70	Thr	Val	Ala	Glu	Ser 75	Ala	Glu	Glu	Glu	Leu 80
55	Gln	Gln	Ala	Gly	Asp 85	Gln	Glu	Leu	Leu	His 90	Gln	Ala	Lys	Asp	Phe 95	Gly
	Asn	Tyr	Leu	Phe 100	Asn	Phe	Ala	Ser	Ala 105	Ala	Thr	Lys	Lys	Ile 110	Thr	Glu
60	Ser	Val	Ala	Glu	Thr	Ala	Gln	Thr	Ile	Lys	Lys	Ser	Val	Glu	Glu	Gly

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			115					120					125			
_	Lys	Ile 130	Asp	Gly	Ile	Ile	Asp 135	Lys	Thr	Ile	Ile	Gly 140	Asp	Phe	Gln	Lys
5	Glu 145	Gln	Lys	Lys	Phe	Val 150	Glu	Glu	Gln	His	Thr 155	Lys	Lys	Ser	Glu	Ala 160
10	Ala	Val	Pro	Pro	Trp 165	Val	Asp	Thr	Asn	Asp 170	Glu	Glu	Thr	Ile	Gln 175	Gln
	Gln	Ile	Leu	Ala 180	Leu	Ser	Ala	Asp	Lys 185	Arg	Asn	Phe	Leu	Arg 190	Asp	Pro
15	Pro	Ala	Gly 195		Gln	Phe	Asn	Phe 200	Asp	Phe	Asp	Gln	Met 205	Tyr	Pro	Val
20	Ala	Leu 210		Met	Leu											
25	(2)	INF		SEQU	ENCE (A) I	E CHA LENG! IYPE	ARACT	NO: TERIS 49 au ino a : li	STICS mino acid		ds					
30		: Arg				ı Val		IPTIO			l Lys			ı Val	. Phe	e Trp
35	Ar	g Ası	n Tyi	r Phe		r Arg	g Vai	l Se	r Let 25		e Ly:	s Glr	n Sei	c Ala		n Leu
	Th	r Al	a Lei		a Ala	a Glı	n Gli	n Gli 4		a Ala	a Gl	y Ly:	s Gly 4!		y Glı	ı Glu
40	Gl:	a														
45	(2) IN						NO:								
50					(A) (B) (D)	TYPI TOP	STH: E: ai OLOG	TERI 76 a mino Y: 1: RIPT	amino ació inear	ac:		NO: 2	247 :			
55	Se	er Th	ır Se	er Pr	ro Gl	Ly Va 5	al Se	er G]	.u Ph		al Se LO	er As	sp Al	a Ph		sp Ala 15
55	Cy	ys As	sn Le		sn G: 20	ln Gl	Lu A	sp Le		g Ly 25	ys G	lu Me	et Gl		ln L ∈ 30	eu Val
60	L	eu A		ys Ly 35	ys G	ln G	lu G		nr A. 40	la V	al L	eu G		lu As 45	sp Se	er Ala

```
Asp Trp Glu Lys Glu Leu Gln Gln Glu Leu Gln Glu Tyr Glu Val Val
                               55
  5
       Thr Glu Ser Glu Lys Arg Asp Glu Asn Trp Asp Lys
                            70
10
       (2) INFORMATION FOR SEQ ID NO: 248:
              (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 62 amino acids
                     (B) TYPE: amino acid
15
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 248:
      Ser Pro Trp Glu Ser Arg Arg Val Glu Ser Lys Ala Thr Ser Ala Arg
20
      Cys Gly Leu Trp Gly Ser Gly Pro Arg Arg Pro Ala Ser Gly Met
      Phe Arg Gly Leu Ser Ser Trp Leu Gly Leu Gln Gln Pro Val Ala Gly
25
      Gly Gly Gln Pro Asn Gly Asp Ala Pro Pro Glu Gln Pro Ser
                               55
30
      (2) INFORMATION FOR SEQ ID NO: 249:
             (i) SEQUENCE CHARACTERISTICS:
35
                     (A) LENGTH: 65 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 249:
40
      Pro Val Ala Gly Gly Gly Gln Pro Asn Gly Asp Ala Pro Pro Glu Gln
        1
                        5
                                        10
      Pro Ser Glu Thr Val Ala Glu Ser Ala Glu Glu Glu Leu Gln Gln Ala
                                       25
45
      Gly Asp Gln Glu Leu Leu His Gln Ala Lys Asp Phe Gly Asn Tyr Leu
                                   40
      Phe Asn Phe Ala Ser Ala Ala Thr Lys Lys Ile Thr Glu Ser Val Ala
50
      Glu
       65
55
      (2) INFORMATION FOR SEQ ID NO: 250:
             (i) SEQUENCE CHARACTERISTICS:
60
                    (A) LENGTH: 72 amino acids
```

	(B) TYPE: amino acid
	(D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 250:
5	Phe Gln Lys Glu Gln Lys Lys Phe Val Glu Glu Gln His Thr Lys Lys 1 5 10 15
	Ser Glu Ala Ala Val Pro Pro Trp Val Asp Thr Asn Asp Glu Glu Thr 20 25 30
10	Ile Gln Gln Ile Leu Ala Leu Ser Ala Asp Lys Arg Asn Phe Leu 35 40 45
15	Arg Asp Pro Pro Ala Gly Val Gln Phe Asn Phe Asp Phe Asp Gln Met 50 55 60
	Tyr Pro Val Ala Leu Val Met Leu 65 70
20	
	(2) INFORMATION FOR SEQ ID NO: 251:
25	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 28 amino acids
	(B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 251:
30	Pro Phe Ile Cys Val Ala Arg Asn Pro Val Ser Arg Asn Phe Ser Ser 1 5 10 15
35	Pro Ile Leu Ala Arg Lys Leu Cys Glu Gly Ala Ala 20 25
	(2) INFORMATION FOR SEQ ID NO: 252:
40	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 33 amino acids(B) TYPE: amino acid
	(D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 252:
45	Lys Glu Asp Pro Ala Asn Thr Val Tyr Ser Thr Val Glu Ile Pro Lys 1 5 10 15
50	Lys Met Glu Asn Pro His Ser Leu Leu Thr Met Pro Asp Thr Pro Arg 20 25 30
	Leu
55	
	(2) INFORMATION FOR SEQ ID NO: 253:
60	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 227 amino acids

			/ <u>-</u> : \		(D) :	TYPE:	OGY:	lir	near							
=						E DE										
5	Ala 1		Ala	Val	Leu 5	Leu	Asp	Leu	Pro	Asn 10	Ser	Gly	Gly	Glu	Ala 15	
10	Ala	Lys	Lys	Leu 20		Asn	Asn	Cys	Val 25	Phe	Ala	Pro	Ala	Asp 30	Val	Thr
	Ser	Glu	Lys 35	Asp	Val	Gln	Thr	Ala 40	Leu	Ala	Leu	Ala	Lys 45	Gly	Lys	Phe
15	Gly	Arg 50	Val	Asp	Val	Ala	Val 55	Asn	Cys	Ala	Gly	Ile 60	Ala	Val	Ala	Ser
	Lys 65	Thr	Tyr	Asn	Leu	Lys 70	Lys	Gly	Gln	Thr	His 75	Thr	Leu	Glu	Asp	Phe 80
20	Gln	Arg	Val	Leu	Asp 85	Val	Asn	Leu	Met	Gly 90	Thr	Phe	Asn	Val	Ile 95	Arg
25	Leu	Val	Ala	Gly 100	Glu	Met	Gly	Gln	Asn 105	Glu	Pro	Asp	Gln	Gly 110	Gly	Gln
-0	Arg	Gly	Val 115	Ile	Ile	Asn	Thr	Ala 120	Ser	Val	Ala	Ala	Phe 125	Glu	Gly	Gln
30	Val	Gly 130	Gln	Ala	Ala	Tyr	Ser 135	Ala	Ser	Lys	Gly	Gly 140	Ile	Val	Gly	Met
	Thr 145	Leu	Pro	Ile	Ala	Arg 150	Asp	Leu	Ala	Pro	Ile 155	Gly	Ile	Arg	Val	Met 160
35	Thr	Ile	Ala	Pro	Gly 165	Leu	Phe	Gly	Thr	Pro 170	Leu	Leu	Thr	Ser	Leu 175	Pro
40	Glu	Lys	Val	Cys 180	Asn	Phe	Leu	Ala	Ser 185	Gln	Val	Pro	Phe	Pro 190	Ser	Arg
10	Leu	Gly	Asp 195	Pro	Ala	Glu	Tyr	Ala 200	His	Leu	Val	Gln	Ala 205	Ile	Ile	Glu
45	Asn	Pro 210	Phe	Leu	Asn	Gly	Glu 215	Val	Ile	Arg	Leu	Asp 220	Gly	Ala	Ile	Arg
	Met 225	Gln	Pro													
50																
	(2)	INFO	RMAT	NOI	FOR	SEQ	ID N	IO: 2	54:							
55			(i) S	()	A) L1 3) T	CHAF ENGTI YPE:	H: 29	am:	ino a		6					
		+	(xi)			OPOLO E DES				EQ II	NO:	254	ł:			
60	Ser	Val	Ala	Ala	Phe	Glu	Gly	Gln	Val	Gly	Gln	Ala	Ala	Tyr	Ser	Ala

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345

	1	5		10	15
5	Ser Lys Gly	Gly Ile Val C	Gly Met Thr 25	Leu Pro Ile	Ala
	(2) INFORMAT	TION FOR SEQ 1	ID NO: 255:		
10		(B) TYPE:	: 61 amino a amino acid GY: linear	acids	5:
15		-			Leu Cys Arg Trp 15
20	Ala Gln Lys	His Lys Asn '	Trp Arg Phe 25	Gln Lys Thr	Arg Gln Thr Trp
	Leu Leu Leu 35	His Met Tyr	Asp Ser Asp 40	Lys Val Pro	Asp Glu His Phe 45
25	Ser Thr Leu 50	Leu Ala Tyr	Leu Glu Gly 55	Leu Gln Gly	
30	(2) INFORMA	TION FOR SEQ	ID NO: 256:		
35		(B) TYPE:	H: 22 amino amino acid OGY: linear	acids	56:
33					ı Ser Gln Phe Tyr 15
40	Ile Asn Lys	: Leu Cys Phe 20			
45		ATION FOR SEQ			
	(i)	(B) TYPE:	H: 22 amino amino acid	acids	٠
50	(xi	(D) TOPOL) SEQUENCE DE	OGY: linear	_	57 :
55	Cys Trp Ile	e Lys Tyr Cys 5	Leu Thr Le	ı Met Gln As 10	n Ala Gln Leu Ser 15
	Met Gln As	p Asn Ile Gly 20			

BNSDOCID: <WO 9842738A1>

```
(2) INFORMATION FOR SEQ ID NO: 258:
              (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 25 amino acids
 5
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 258:
      Lys Val Ser Tyr Leu Arg Pro Leu Asp Phe Glu Glu Ala Arg Glu Leu
10
      Phe Leu Leu Gly Gln His Tyr Val Phe
                   20
15
      (2) INFORMATION FOR SEQ ID NO: 259:
              (i) SEQUENCE CHARACTERISTICS:
20
                     (A) LENGTH: 25 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 259:
25
      Met Glu Arg Arg Cys Lys Met His Lys Arg Xaa Ile Ala Met Leu Glu
                        5
      Pro Leu Thr Val Asp Leu Asn Pro Gln
                   20
30
      (2) INFORMATION FOR SEQ ID NO: 260:
35
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 23 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 260:
40
      Ser His Ile Val Lys Lys Ile Asn Asn Leu Asn Lys Ser Ala Leu Lys
      Tyr Tyr Gln Leu Phe Leu Asp
45
                   20
      (2) INFORMATION FOR SEQ ID NO: 261:
50
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 64 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
55
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 261:
      Phe Thr His Leu Ser Thr Cys Leu Leu Ser Leu Leu Leu Val Arg Met
                                           10
60
      Ser Gly Phe Leu Leu Leu Ala Arg Ala Ser Pro Ser Ile Cys Ala Leu
```

	20	25	30
	Asp Ser Ser Cys Phe Val Gl	n Glu Tyr Cys Ser 40	Ser Tyr Ser Ser Ser 45
5	Cys Phe Leu His Gln His Ph 50 5	e Pro Ser Leu Leu 5	Asp His Leu Cys Gln 60
10			
15	(2) INFORMATION FOR SEQ ID		
20	(A) LENGTH: (B) TYPE: au (D) TOPOLOG (xi) SEQUENCE DESCI	Y: linear	v: 262:
	Phe Leu Leu Leu Ala Arg Al 1 5	la Ser Pro Ser Ile 10	Cys Ala Leu Asp Ser 15
25	Ser Cys Phe Val Gln Glu Ty 20	γr	
30	(2) INFORMATION FOR SEQ I		
35	(B) TYPE: a (D) TOPOLOG	53 amino acids mino acid	D: 2 63:
40	Pro Asp Gly Arg Val Thr A 1 5 Phe Gly Met Ile Gly Leu L	10	15
45	20 Pro Gly Met Val His Leu A 35	25	30
	Leu Asn Leu Asn Ser 50		
50			
	(2) INFORMATION FOR SEQ :		
55	(B) TYPE: (D) TOPOLO	ACTERISTICS: 1: 41 amino acids amino acid XGY: linear CRIPTION: SEQ ID N	vio - 264 -
60	Glu Asp Leu Leu Phe Tyr		

	1				5	i				10					15	
5	Gln	Leu	Leu	Ala 20		Val	Glu	Leu	Phe 25		Arg	Asp	Trp	Arg 30	Tyr	His
J	Lys	Glu	Glu 35		Val	Trp	Ile	Thr 40								
10	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	265:							
15				•	(A) I (B) T (D) T	CHA LENGT TYPE: TOPOI TE DE	TH: 2 ami .OGY:	4 an no a lir	nino cid near	acid): 26	5:			
20	Val 1	His	Leu	Ala	Leu 5	Gly	Ser	Asp	Leu	Thr 10	Thr	Leu	Gly	Leu	Asn 15	Leu
	Asn	Ser	Pro	Glu 20	Asn	Leu	Tyr	Pro								
25																
	(2)	INF	ORMA'	rion	FOR	SEQ	ID I	NO:	266:							
30				((A) L B) T D) T	CHA ENGT YPE: OPOL E DE	H: 4 ami OGY:	1 am no a lin	ino cid ear	acid		: 26	6:			
35	His 1					Pro										
40	(2)	INF	ORMA:	MOIT	FOR	SEQ	ID I	10: 2	267 :							
45				(A) L B) T D) T	CHAI ENGT YPE: OPOL E DE	H: 7 ami OGY:	5 am no a lin	ino cid ear	acid		: 26	7:			
50	Gly 1	Arg	Ile	Ile	Asp 5	Thr	Ser	Leu	Thr	Arg 10	Asp	Pro	Leu	Val	Ile 15	Glu
	Leu	Gly	Gln	Lys 20	Gln	Val	Ile	Pro	Gly 25	Leu	Glu	Gln	Ser	Leu 30	Leu	Asp
55	Met	Cys	Val 35	Gly	Glu	Lys	Arg	Arg 40	Ala	Ile	Ile	Pro	Ser 45	His	Leu	Ala
	Tyr	Gly 50	Lys	Arg	Gly	Phe	Pro 55	Pro	Ser	Val	Pro	Ala 60	Asp	Ala	Val	Val
60	Gln	Tyr	Asp	Val	Glu	Leu	Ile	Ala	Leu	Ile	Arg					

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349 75 70 65 5 (2) INFORMATION FOR SEQ ID NO: 268: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid 10 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 268: Ile His Tyr Thr Gly Ser Leu Val Asp Gly Arg Ile Ile Asp Thr Ser 5 15 20 (2) INFORMATION FOR SEQ ID NO: 269: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 20 amino acids 25 (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 269: Cys Glu Ser Pro Glu Ser Pro Ala Gln Pro Ser Gly Ser Ser Leu Pro 30 5 Ala Trp Tyr His 20 35 (2) INFORMATION FOR SEQ ID NO: 270: (i) SEQUENCE CHARACTERISTICS: 40 (A) LENGTH: 95 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 270: Glu Glu Ala Gly Ala Gly Arg Arg Cys Ser His Gly Gly Ala Arg Pro 45 10 Ala Gly Leu Gly Asn Glu Gly Leu Gly Leu Gly Gly Asp Pro Asp His 25 20 50 Thr Asp Thr Gly Ser Arg Ser Lys Gln Arg Ile Asn Asn Trp Lys Glu Ser Lys His Lys Val Ile Met Ala Ser Ala Ser Ala Arg Gly Asn Gln 55 50 Asp Lys Asp Ala His Phe Pro Pro Pro Ser Lys Gln Ser Leu Leu Phe

70

Cys Pro Lys Ser Lys Leu His Ile His Arg Ala Glu Ile Ser Lys

85 90 95 5 (2) INFORMATION FOR SEQ ID NO: 271: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 amino acids (B) TYPE: amino acid 10 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 271: Ser Lys Gln Arg Ile Asn Asn Trp Lys Glu Ser Lys His Lys Val Ile 15 Met Ala Ser Ala Ser Ala Arg 20 20 (2) INFORMATION FOR SEQ ID NO: 272: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 32 amino acids 25 (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 272: Leu Phe His Trp Ala Cys Leu Asn Glu Arg Ala Ala Gln Leu Pro Arg 30 Asn Thr Ala Xaa Ala Gly Tyr Gln Cys Pro Ser Cys Asn Gly Pro Ser 25 35 40 (2) INFORMATION FOR SEQ ID NO: 273: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 185 amino acids (B) TYPE: amino acid 45 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 273: Phe Tyr Ile Tyr Tyr Arg Pro Thr Asp Ser Asp Asn Asp Ser Asp Tyr 5 50 Lys Lys Asp Met Val Glu Gly Asp Lys Tyr Trp His Ser Ile Ser His 25 Leu Gln Pro Glu Thr Ser Tyr Asp Ile Lys Met Gln Cys Phe Asn Glu 55 40 Gly Gly Glu Ser Glu Phe Ser Asn Val Met Ile Cys Glu Thr Lys Ala

Arg Lys Ser Ser Gly Gln Pro Gly Arg Leu Pro Pro Pro Thr Leu Ala

	65					70					75					80
	Pro	Pro	Gln	Pro	Pro 85	Leu	Pro	Glu	Thr	Ile 90	Glu	Arg	Pro	Val	Gly 95	Thr
5	Gly	Ala	Met	Val 100	Ala	Arg	Ser	Ser	Asp 105	Leu	Pro	Tyr	Leu	Ile 110	Val	Gly
10	Val	Val	Leu 115	Gly	Ser	Ile	Val	Leu 120	Ile	Ile	Val	Thr	Phe 125	Ile	Pro	Phe
	Cys	Leu 130	Trp	Arg	Ala	Trp	Ser 135	Lys	Gln	Lys	His	Thr 140	Thr	Asp	Leu	Gly
15	Phe 145	Pro	Arg	Ser	Ala	Leu 150	Pro	Pro	Ser	Cys	Pro 155	Tyr	Thr	Met	Val	Pro 160
20	Leu	Gly	Gly	Leu	Pro 165	Gly	His	Gln	Ala	Val 170		Ser	Pro	Thr	Ser 175	
20	Ala	Ser	Val	Asp 180	Gly	Pro	Val	Leu	Met 185							
25	(2)	INF	FORMA	MOIT	FOR	: SEQ) ID	NO:	274:							
				SEQU	JENCE	E CHA	ARAC'	reris		S:	ds					
30			(xi		(D)	ropo:	LOGY	ino a : li: IPTI		SEQ	ID N	o: 2	74:			
35	Tyr 1		е Тул	с Туг		g Pro	o Thi	r Ası	Sei	Ası		n Asp) Sei	c Asp	тул 19	Lys
	Lys	s As	p Me	t Val		ı Gly	y As	p Ly:	s Ty:		p Hi:	s Sei	c Il	e Sei		s Leu
40	Glı	n Pr	o Gl		r Se:	r Ty:	r As	p Il.		s Me	t Gl	n Cy	s Ph 4		n Gl	u Gly
45			0	r Gl	u Ph	e Se		n Va 5	l Me	t Il	e Cy	s Gl	u Th O	r Ly	s Al	a Arg
	. 6															
50	(2) IN	VFORM	IATIC	N FC	R SE	Q II	NO:	275	:						
55) SE((A) (B) (D)	TYP:	GTH: E: a OLOG	30 a mino Y: 1	amino acio inea:	o ac d r		NO: 3	275 :			
60	As	sn V	al A	rg A	la Le	eu Le S	eu H	is A	rg Me		ro G: 10	lu Pi	ro P	ro Ly		le Asn 15

	1111	. Alc	r rys	20		ASI	ASI	Lys	Arg 25		Asn	Leu	Ser	Leu 30		
5																
	(2)	INF	FORMA	TION	FOR	SEQ	ID	NO:	276:							
10				((A) I (B) T (D) T	ENGT YPE : YPOI	TH: 1 ami OGY:	ERIS 185 a ino a lin	mino cid ear	aci		: 27	6:			
15	Asn 1	Thr												Phe	Gln 15	Pro
20	Phe	Ala	Leu	Asn 20	His	Gln	Lys	Asp	Ile 25	Gln	Val	Leu	Met	Gly 30	Ser	Leu
	Val	Tyr	Leu 35	Arg	Gln	Gly	Ile	Glu 40	Asn	Ser	Pro	Tyr	Val 45	His	Leu	Let
25	Asp	Ala 50	Asn	Gln	Trp	Ala	Asp 55	Ile	Cys	Asp	Ile	Phe 60	Thr	Arg	Asp	Ala
	Cys 65	Ala	Leu	Leu	Gly	Leu 70	Ser	Val	Glu	Ser	Pro 75	Leu	Ser	Val	Ser	Phe 80
30	Ser	Ala	Gly	Cys	Val 85	Ala	Leu	Pro	Ala	Leu 90	Ile	Asn	Ile	Lys	Ala 95	Val
35	Ile	Glu	Gln	Arg 100	Gln	Cys	Thr	Gly	Val 105	Trp	Asn	Gln	Lys	Asp 110	Glu	Leu
	Pro	Ile	Glu 115	Val	Asp	Leu	Gly	Lys 120	Lys	Cys	Trp	Tyr	His 125	Ser	Ile	Phe
40	Ala	Cys 130	Pro	Ile	Leu	Arg	Gln 135	Gln	Thr	Thr	Asp	Asn 140	Asn	Pro	Pro	Met
	Lys 145	Leu	Val	Cys	Gly	His 150	Ile	Ile	Ser	Arg	Asp 155	Ala	Leu	Asn	Lys	Met 160
45	Phe	Asn	Gly	Ser	Lys 165	Leu	Lys	Cys	Pro	Tyr 170	Cys	Pro	Met	Glu	Gln 175	Ser
50	Pro	Gly	Asp	Ala 180	Lys	Gln	Ile	Phe	Phe 185							
	(2)	INF	ORMAT	CION	FOR	SEQ	ID 1	IO: 2	77:							
55			(i) s	() ()	A) Li B) T	ENGTI YPE :	H: 6	ERIST 5 am no ao line	ino a		5					
60			(xi)							EQ II	NO:	277	7:			

	Ser 1	Tyr	Leu	Ser	Ala 5	Cys	Phe	Ala	Gly	Cys 10	Asn	Ser	Thr	Asn	Leu 15	unr
5	Gly	Cys	Ala	Cys 20	Leu	Thr	Thr	Val	Pro 25	Ala	Glu	Asn	Ala	Thr 30	Val	Val
	Pro	Gly	Lys 35	Cys	Pro	Ser	Pro	Gly 40	Cys	Gln	Glu	Ala	Phe 45	Leu	Thr	Phe
10	Leu	Cys 50	Val	Met	Cys	Ile	Cys 55		Leu	Ile	Gly	Ala 60	Met	Ala	Arg	His
15	Pro 65															
20	(2)	INF	(i)		ENCE (A) I (B) 1	CHA LENGI TYPE:	RACT TH: { : am: LOGY	TERIS 34 ar ino a : lir	TICS mino acid near	acio		o: 27	78:			
25	Pro 1			l Il∈		. Leu					. Ser			ı Leu	Lys 15	Ser
30	Тут	Ala	ı Lev	ı Gly 20		. Leu	ı Phe	e Leu	ı Lev 25		ı Arç	g Lev	ı Lev	Gly 30		Ile
	Pro	Pro	Pro 39		ı Ile	e Phe	e Gly	y Ala 40		/ Ile	e Ası) Ser	Thu 45		: Leu	Phe
35	Tr	Sei 50		r Phe	e Cys	s Gly	/ Gl: 5!		n Gly	y Ala	a Cy:	s Va:		1 Туг	Asp) Asn
40	6	5		r Arg		r Lei 7		r Va	l Se:	r Il	e Al: 7		e Ala	a Lev	ı Lys	Ser 80
45	(2) IN		ATIO												
50				SEQ	(A) (B) (D)	TYPI TOPO	STH: E: ar OLOG	182 mino Y: 1:	amin acio inean	no ad 1 r		NO: 2	279:			
55	G1	ln Se	er Le	eu Ph	ne Th	ır Ar 5	g Pł	ne Va	al Ar		al G1 10	ly Va	al Pr	o Th		l Asp 5
	Le	eu As	sp A		ln G] 20	ly Ai	g A	la Ar		la Se 25	er Le	eu Cy	/s Xā		а Ту Ю	r Asn
60	T	rp Ai	rg T	yr Ly	ys As	sn Le	eu G	ly As	sn Le	eu P	ro H	is Vá	al G	ln Le	eu Le	u Pro

			35					40					45			
5	Glu	Phe 50	Ser	Thr	Ala	Asn	Ala 55		Leu	Leu	Tyr	Asp 60	Phe	Gln	Leu	Ile
J	Asn 65	Val	Glu	Asp	Phe	Gln 70	Gly	Val	Gly	Glu	Ser 75	Glu	Pro	Asn	Pro	Туг 80
10	Phe	Tyr	Gln	Asn	Leu 85	Gly	Glu	Ala	Glu	Tyr 90	Val	Val	Ala	Leu	Phe 95	Met
	Tyr	Met	Cys	Leu 100	Leu	Gly	Tyr	Pro	Ala 105	Asp	Lys	Ile	Ser	Ile 110	Leu	Thr
15	Thr	Tyr	Asn 115	Gly	Gln	Lys	His	Leu 120	Ile	Arg	Asp	Ile	Ile 125	Asn	Arg	Arg
20	Cys	Gly 130	Asn	Asn	Pro	Leu	Ile 135	Gly	Arg	Pro	Asn	Lys 140	Val	Thr	Thr	Val
	Asp 145	Arg	Phe	Gln	Gly	Gln 150	Gln	Asn	Asp	Tyr	Ile 155	Leu	Leu	Ser	Leu	Val 160
25	Arg	Thr	Arg	Ala	Val 165	Gly	His	Leu	Arg	Asp 170	Val	Arg	Arg	Leu	Val 175	Val
	Ala	Met	Ser	Arg 180	Ala	Arg										
30				•												
	(2)							NO: 2								
35				(. (: (:	A) L: B) T D) T	ENGT: YPE : OPOL	H: 7 ami: OGY:	ERIST 7 am no a lin	ino a cid ear	acid						
40	Y		(xi)					PTIO								
40	Leu 1	vai	ьуs	Glu	Ala 5	Lys	Ile	Ile	Ala	Met 10	Thr	Cys	Thr	His	Ala 15	Ala
45	Leu	Lys	Arg	His 20	Asp	Leu	Val	Lys	Leu 25	Gly	Phe	Lys	Tyr	Asp 30	Asn	Ile
	Leu	Met	Glu 35	Glu	Ala	Ala	Gln	Ile 40	Leu	Glu	Ile	Glu	Thr 45	Phe	Ile	Pro
50	Leu	Leu 50	Leu	Gln	Asn	Pro	Gln 55	Asp	Gly	Phe	Ser	Arg 60	Leu	Lys	Arg	Trp
<i></i>	Ile 65	Met	Ile	Gly	Asp	His 70	His	Gln	Leu	Pro	Pro 75	Val	Ile			
55	(2)	TNEC	\D\#\ \ #	TON	707											
	(2)					-		10: 2								
60			(1) 2					ERIST 25 ar			ds					

						YPE: OPOLO										
			(xi)	SEQU						EQ II	NO:	281	.:			
5	Asp 1	Thr	Tyr	Pro	Asn 5	Glu	Glu	Lys	Gln	Gln 10	Glu	Arg	Val	Phe	Pro 15	Xaa
	Xaa	Ser	Ala	Met 20	Val	Asn	Asn	Gly	Ser 25	Leu	Ser	Tyr	Asp	His 30	Glu	Arg
10	Asp	Gly	Arg 35	Pro	Thr	Glu	Leu	Gly 40	Gly	Cys	Xaa	Ala	Ile 45	Val	Arg	Asn
15	Leu	His 50	Tyr	Asp	Thr	Phe	Leu 55	Val	Ile	Arg	Tyr	Val 60	Lys	Arg	His	Leu
	Thr 65	Ile	Met	Met	Asp	Ile 70	Asp	Gly	Lys	His	Glu 75	Trp	Arg	Asp	Cys	Ile 80
20	Glu	Val	Pro	Gly	Val 85	Arg	Leu	Pro	Arg	Gly 90	Tyr	Tyr	Phe	Gly	Thr 95	Ser
25	Ser	Ile	Thr	Gly 100	Asp	Leu	Ser	Asp	Asn 105	His	Asp	Val	Ile	Ser 110	Leu	Lys
25	Leu	Phe	Glu 115	Leu	Thr	Val	Glu	Arg 120		Pro	Glu	Glu	Glu 125			
30	(2)	TATE	· ADM »	TION	FOE	SEO	TD	N∩ ·	282.							
	(2)	11/1		SEOU						S:						
35			, -,	((A) : (B) '	LENGT TYPE: TOPOI	TH: 8	35 ar ino a	mino acid		ds					
			(xi)	SEÇ	QUEN(CE DE	SCR	PTI	ON: S	SEQ :	ID NO): 28	32:			
40	Leu 1		s Arg	g Glu		s Ser	. Len	sei	Lys	Pro 10		Glr	ı Gly	/ Val	. Gly 15	Thr
	Gly	/ Se	r Sei	Ser 20		ı Trţ	Ası	ı Leı	ı Met		/ Asr	n Ala	a Me	: Val		Thr
45	Glr	ту:	r Ile 3!		g Le	u Thi	Pro	As ₁		: Gli	n Sei	: Ly:	s Gl:		/ Ala	a Leu
50	Tr	As 5		g Val	l Pr	o Cys	s Phe		u Arg	g As	p Trj	9 Gli 6		u Gli	n Vai	l His
50	Pho		s Il	e His	s Gl	y Gla 7		y Ly	s Ly:	s As	n Lei 7		s Gl	y As	o Gl	y Leu 80
55	Al	a Il	e Tr	р Туг		r 5										
60	(2) IN	IFORM	OITA	N FC	R SE	Q ID	NO:	283	:						

```
(i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 32 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
  5
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 283:
       Pro Gly Thr Leu Gln Cys Ser Ala Leu His His Asp Pro Gly Cys Ala
                        5
                                           10
10
      Asn Cys Ser Arg Phe Cys Arg Asp Cys Ser Pro Pro Ala Cys Gln Cys
                                       25
15
       (2) INFORMATION FOR SEQ ID NO: 284:
20
              (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 27 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 284:
25
      Phe Leu Tyr Asp Val Leu Met Xaa His Glu Ala Val Met Arg Thr His
               5
                                   10
      Gln Ile Gln Leu Pro Asp Pro Glu Phe Pro Ser
30
                                      25
      (2) INFORMATION FOR SEQ ID NO: 285:
35
              (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 6 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
40
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 285:
      Gly Trp Tyr Trp Cys Gly
        1
45
      (2) INFORMATION FOR SEQ ID NO: 286:
              (i) SEQUENCE CHARACTERISTICS:
50
                     (A) LENGTH: 129 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 286:
55
      Met Lys Val Gly Ala Arg Ile Arg Val Lys Met Ser Val Asn Lys Ala
      His Pro Val Val Ser Thr His Trp Arg Trp Pro Ala Glu Trp Pro Gln
                                  25
                   20
60
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	Met	Phe	Leu 35	His	Leu	Ala	Gln	Glu 40	Pro	Arg	Thr	Glu	Val 45	Lys	Ser	Arg
5	Pro	Leu 50	Gly	Leu	Ala	Gly	Phe 55	Ile	Arg	Gln	Asp	Ser 60	Lys	Thr	Arg	Lys
	Pro 65	Leu	Glu	Gln	Glu	Thr 70	Ile	Met	Ser	Ala	Ala 75	Asp	Thr	Ala	Leu	Trp 80
10	Pro	Tyr	Gly	His	Gly 85	Asn	Arg	Glu	His	Gln 90	Glu	Asn	Glu	Leu	Gln 95	Lys
15	Tyr	Leu	Gln	Tyr 100		Asp	Met	His	Leu 105	Leu	Asp	Ser	Gly	Gln 110	Ser	Leu
13	Gly	His	Thr 115		Thr	Leu	Gln	Gly 120		His	Asn	Leu	Thr 125	Ala	Leu	Asn
20	Ile															
25	(2)	INF		ATION SEQU	JENCI	E CHA	ARAC'		STICS	5:	ds					
30) SEG	(B) (D) QUEN	TYPE TOPO	: am LOGY ESCR	ino : li IPTI	acid near ON:	SEQ	ID N					
		Lei l	ı Hi:	s Lys		n Ser 5	r Va	l Se:	r Gli	n Ile 10		r Vai	l Lei	ı Sei	r Gly	y Gly 5
35				2	0				2	5				3	0	y Met
40	Se:			p As	p Va	l Ly	s Se	r Le 4		u Se	r Al	a Le	u Ly 4		p Le	u Lys
45	(2) IN	IFORM	OITAI	N FC	R SE	Q II	NO:	288	:						
50) SE((A) (B) (D)	TYP!	GTH: E: a: OLOG	21 a mino Y: 1	amino acio inea:	o ac d r		NO: 1	288:			
55	G]	lu Al	la S	er Ly	ys Se	er Se 5	er H	is A	la Gi		eu As 10	sp Le	eu Pl	ne Se	er Va	al Ala 15
	A.	la C	ys H	is A	rg Pl 20	he										
60																

	(2) INFORMATION FOR SEQ ID NO: 289:
5	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 21 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 289:
10	Tyr Met Gly Lys Gly Ser Met Thr Gly Leu Ala Leu Lys His Met Phe 1 5 10 15
15	Glu Arg Ser Phe Thr 20
20	(2) INFORMATION FOR SEQ ID NO: 290: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 27 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 290: Val Thr Gly Ile Ile Asp Ser Leu Thr Ile Ser Pro Lys Ala Ala Arg 1 5 10 15
30	Val Gly Leu Leu Gln Tyr Ser Thr Gln Val His 20 25
35 40	(2) INFORMATION FOR SEQ ID NO: 291: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 24 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 291:
	Thr Glu Phe Thr Leu Arg Asn Phe Asn Ser Ala Lys Asp Met Lys Lys 1 5 10 15
4 5	Ala Val Ala His Met Lys Tyr Met 20
50	(2) INFORMATION FOR SEQ ID NO: 292: (i) SEQUENCE CHARACTERISTICS:
55	(A) LENGTH: 27 amino acids(B) TYPE: amino acid(D) TOPOLOGY: linear(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 292:
50	Gly Lys Gly Ser Met Thr Gly Leu Ala Leu Lys His Met Phe Glu Arg 1 5 10 15

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Ser Phe Thr Gln Gly Glu Gly Ala Arg Pro Phe 20 25

5 (2) INFORMATION FOR SEQ ID NO: 293:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 44 amino acids
 - (B) TYPE: amino acid
 - (D) TOPOLOGY: linear
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 293:
- Ser Thr Arg Val Pro Arg Ala Ala Ile Val Phe Thr Asp Gly Arg Ala
 15 1 5 10 15

Gln Asp Asp Val Ser Glu Trp Ala Ser Lys Ala Lys Ala Asn Gly Ile 20 25 30

- 20 Thr Met Tyr Ala Val Gly Val Gly Lys Ala Ile Glu
- 25 (2) INFORMATION FOR SEQ ID NO: 294:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 42 amino acids
 - (B) TYPE: amino acid
- 30 (D) TOPOLOGY: linear
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 294:

Glu Glu Leu Gln Glu Ile Ala Ser Glu Pro Thr Asn Lys His Leu Phe 1 5 10 15

Tyr Ala Glu Asp Phe Ser Thr Met Asp Glu Ile Ser Glu Lys Leu Lys
20 25 30

Lys Gly Ile Cys Glu Ala Leu Glu Asp Ser 40 35 40

- (2) INFORMATION FOR SEQ ID NO: 295:
- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 11 amino acids
 - (B) TYPE: amino acid
 - (D) TOPOLOGY: linear
- 50 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 295:

Thr Gln Arg Leu Glu Glu Met Thr Gln Arg Met
1 5 10

55

45

- (2) INFORMATION FOR SEQ ID NO: 296:
- (i) SEQUENCE CHARACTERISTICS:
- 60 (A) LENGTH: 10 amino acids

```
(B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 296:
 5
      Pro Gln Gly Cys Pro Glu Gln Pro Leu His
10
      (2) INFORMATION FOR SEQ ID NO: 297:
              (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 33 amino acids
                     (B) TYPE: amino acid
15
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 297:
      Arg Cys Lys Lys Cys Thr Glu Gly Pro Ile Asp Leu Val Phe Val Ile
                                           10
20
      Asp Gly Ser Lys Ser Leu Gly Glu Glu Asn Phe Glu Val Val Lys Gln
      Phe
25
      (2) INFORMATION FOR SEQ ID NO: 298:
30
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 60 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
35
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 298:
      Met Ala Ala Leu Leu Arg His Val Gly Arg His Cys Leu Arg Ala
40
      His Phe Ser Pro Gln Leu Cys Ile Arg Asn Ala Val Pro Leu Gly Thr
                   20
                                       25
      Thr Ala Lys Glu Glu Met Glu Arg Phe Trp Asn Lys Asn Ile Gly Ser
45
      Asn Arg Pro Leu Ser Pro His Ile Thr Ile Tyr Ser
           50
50
      (2) INFORMATION FOR SEQ ID NO: 299:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 32 amino acids
55
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 299:
     Val Phe Pro Leu Met Tyr His Thr Trp Asn Gly Ile Arg His Leu Met
60
                       5
```

	Trp Asp Leu Gly Lys Gly Leu Lys Ile Pro Gln Leu Tyr Gln Ser Gly 20 25 30
5	
10	(2) INFORMATION FOR SEQ ID NO: 300:
15	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 17 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 300:
20	Met Ala Ala Leu Leu Leu Arg His Val Gly Arg His Cys Leu Arg Ala 1 5 10 15 His
25	(2) INFORMATION FOR SEQ ID NO: 301:
30	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 18 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 301:
35	Val Lys Ser Leu Cys Leu Gly Pro Ala Leu Ile His Thr Ala Lys Phe 1 5 10 15
	Ala Leu
40	202
45	(2) INFORMATION FOR SEQ ID NO: 302: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 302:
50	Val Phe Pro Leu Met Tyr His Thr Trp Asn Gly Ile Arg His Leu Met 1 5 10 15
55	Trp Asp Leu Gly Lys Gly Leu 20
	(2) INFORMATION FOR SEQ ID NO: 303:
60	(i) SEQUENCE CHARACTERISTICS:

```
(A) LENGTH: 22 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 303:
 5
      Arg Val Trp Asp Val Arg Pro Phe Ala Pro Lys Glu Arg Cys Val Lys
      Ile Phe Gln Gly Asn Val
10
                   20
      (2) INFORMATION FOR SEQ ID NO: 304:
15
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 30 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
20
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 304:
      His Asn Phe Glu Lys Asn Leu Leu Arg Cys Ser Trp Ser Pro Asp Gly
25
      Ser Lys Ile Ala Ala Gly Ser Ala Asp Arg Phe Val Tyr Val
                                      25
                   20
30
      (2) INFORMATION FOR SEQ ID NO: 305:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 30 amino acids
                     (B) TYPE: amino acid
35
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 305:
      Trp Asp Thr Thr Ser Arg Arg Ile Leu Tyr Lys Leu Pro Gly His Ala
                                         10
40
      Gly Ser Ile Asn Glu Val Ala Phe His Pro Asp Glu Pro Ile
                   20
                                       25
45
      (2) INFORMATION FOR SEQ ID NO: 306:
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 20 amino acids
50
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 306:
      Val Arg Gly Arg Thr Val Leu Arg Pro Gly Leu Asp Ala Glu Pro Glu
55
                        5
                                           10
      Leu Ser Pro Glu
60
```

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(2) INFORMATION FOR SEQ ID NO: 307:
             (i) SEQUENCE CHARACTERISTICS:
 5
                    (A) LENGTH: 19 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 307:
      Glu Gln Arg Val Leu Glu Arg Lys Leu Lys Lys Glu Arg Lys Lys Glu
10
                                          10
      Glu Arg Gln
15
      (2) INFORMATION FOR SEQ ID NO: 308:
20
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 13 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 308:
25
      Arg Leu Arg Glu Ala Gly Leu Val Ala Gln His Pro Pro
                        5
30
       (2) INFORMATION FOR SEQ ID NO: 309:
              (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 17 amino acids
35
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 309:
       Gly Arg Ile Pro Ala Pro Ala Pro Ser Val Pro Ala Gly Pro Asp Ser
 40
                        5
                                            10
       Arg
 45
       (2) INFORMATION FOR SEQ ID NO: 310:
              (i) SEQUENCE CHARACTERISTICS:
 50
                      (A) LENGTH: 42 amino acids
                      (B) TYPE: amino acid
                      (D) TOPOLOGY: linear
               (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 310:
       Thr Gly Cys Val Leu Val Leu Ser Arg Asn Phe Val Gln Tyr Ala Cys
 55
                                             10
       Phe Gly Leu Phe Gly Ile Ile Ala Leu Gln Thr Ile Ala Tyr Ser Ile
                                         25
                    20
 60
```

Leu Trp Asp Leu Lys Phe Leu Met Arg Asn

```
35
 5
      (2) INFORMATION FOR SEQ ID NO: 311:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 55 amino acids
10
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 311:
      Ser Arg Ser Glu Gly Lys Ser Met Phe Ala Gly Val Pro Thr Met Arg
15
                                          10
     Glu Ser Ser Pro Lys Gln Tyr Met Gln Leu Gly Gly Arg Val Leu Leu
20
     Val Leu Met Phe Met Thr Leu Leu His Phe Asp Ala Ser Phe Phe Ser
                                  40
      Ile Val Gln Asn Ile Val Gly
25
      (2) INFORMATION FOR SEQ ID NO: 312:
30
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 60 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 312:
35
     Gly Thr Ala Glu Asp Phe Ala Asp Gln Phe Leu Arg Val Thr Lys Gln
     Tyr Leu Pro His Val Ala Arg Leu Cys Leu Ile Ser Thr Phe Leu Glu
40
     Asp Gly Ile Arg Met Trp Phe Gln Trp Ser Glu Gln Arg Asp Tyr Ile
45
     Asp Thr Thr Trp Asn Cys Gly Tyr Leu Leu Ala Ser
                              55
50
      (2) INFORMATION FOR SEQ ID NO: 313:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 17 amino acids
                    (B) TYPE: amino acid
55
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 313:
     Ala Ser Phe Leu Leu Ser Arg Thr Ser Trp Gly Thr Ala Leu Met Ile
       1 5
60
```

Leu

```
5
     (2) INFORMATION FOR SEQ ID NO: 314:
            (i) SEQUENCE CHARACTERISTICS:
                   (A) LENGTH: 8 amino acids
10
                   (B) TYPE: amino acid
                   (D) TOPOLOGY: linear
            (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 314:
     Leu Met Arg Asn Glu Ser Arg Ser
15
                     5
      1
      (2) INFORMATION FOR SEQ ID NO: 315:
20
             (i) SEQUENCE CHARACTERISTICS:
                   (A) LENGTH: 13 amino acids
                   (B) TYPE: amino acid
                   (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 315:
25
      Ala Ser Phe Leu Leu Ser Arg Thr Ser Trp Gly Thr Ala
       1 5
                                        10
30
      (2) INFORMATION FOR SEQ ID NO: 316:
             (i) SEQUENCE CHARACTERISTICS:
35
                   (A) LENGTH: 20 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 316:
      Phe Ile Ser Phe Ala Asn Ser Arg Ser Ser Glu Asp Thr Lys Gln Met
40
            5
                             10
      Met Ser Ser Phe
45
       (2) INFORMATION FOR SEQ ID NO: 317:
 50
              (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 27 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 317:
 55
       Asp Pro Arg Arg Pro Asn Lys Val Leu Arg Tyr Lys Pro Pro Pro Ser
                  5
        1
       Glu Cys Asn Pro Ala Leu Asp Asp Pro Thr Pro
 60
                   20
```

```
(2) INFORMATION FOR SEQ ID NO: 318:
 5
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 30 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
10
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 318:
      Asp Tyr Met Asn Leu Leu Gly Met Ile Phe Ser Met Cys Gly Leu Met
15
      Leu Lys Leu Lys Trp Cys Ala Trp Val Ala Val Tyr Cys Ser
                   20
                                      25
20
      (2) INFORMATION FOR SEQ ID NO: 319:
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 22 amino acids
                     (B) TYPE: amino acid
25
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 319:
      Met Leu Ser Ile Ser Ala Val Val Met Ser Tyr Leu Gln Asn Pro Gln
                                          10
30
      Pro Met Thr Pro Pro Trp
                   20
35
      (2) INFORMATION FOR SEQ ID NO: 320:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 52 amino acids
40
                     (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 320:
      Ala Ala Gly Asp Gly Asp Val Lys Leu Gly Thr Leu Gly Ser Gly Ser
45
                       5
                                           10
      Glu Ser Ser Asn Asp Gly Gly Ser Glu Ser Pro Gly Asp Ala Gly Ala
50
      Ala Ala Xaa Gly Gly Gly Trp Ala Ala Ala Ala Leu Ala Leu Leu Thr
      Gly Gly Glu
          50
55
      (2) INFORMATION FOR SEQ ID NO: 321:
60
            (i) SEQUENCE CHARACTERISTICS:
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				(1	3) T	ENGTH YPE:	amir	no ao	cid	acio	ds					
_			(xi)			OPOLO E DES				EQ II	ONO:	321	L:			
5	Ala 1	Ala	Asp	Asn	Tyr 5	Gly	Ile	Pro	Arg	Ala 10	Cys	Arg	Asn	Ser	Ala 15	Arg
10	Ser	Tyr	Gly	Ala 20	Ala	Trp	Leu	Leu	Leu 25	Xaa	Pro	Ala	Gly	Ser 30	Ser	Arg
	Val	Glu	Pro 35	Thr	Gln	Asp	Ile	Ser 40	Ile	Ser	Asp	Gln	Leu 45	Gly	Gly	Gln
15	Asp	Val 50	Pro	Val	Phe	Arg	Asn 55	Leu	Ser	Leu	Leu	Val 60	Val	Gly	Val	Gly
20	Ala 65	Val	Phe	Ser	Leu	Leu 70	Phe	His	Leu	Gly	Thr 75	Arg	Glu	Arg	Arg	Arg 80
	Pro	His	Ala	Xaa	Glu 85	Pro	Gly	Glu	His	Thr 90	Pro	Leu	Leu	Ala	Pro 95	Ala
25	Thr	Ala	Gln	Pro 100	Leu	Leu	Leu	Trp	Lys 105		Trp	Leu	Arg	Glu 110	Xaa	Ala
	Phe	Tyr	Gln 115		Gly	Ile	Leu	Tyr 120		Thr	Thr	Arg	Leu 125	Ile	Val	Asn
30	Leu	Ser 130		Thr	Туг	Met	Ala 135		Tyr	Leu	Thr	Tyr 140		Leu	His	Leu
35	Pro 145		. Lys	: Phe	lle	150		Ile	Pro	Leu	155		Tyr	Leu	Ser	Gly 160
	Ph∈	e Lev	ı Ser	: Ser	Phe 165		Met	. Lys	Pro	170		ı Lys	Cys	Ile	: Gly 175	Arg
40	Asr	1														
	(2)) IN	FORM	1OITA	1 FOI	R SEQ) ID	NO:	322	:						
45			(i)	SEQ	(A) (B)	E CHA LENG TYPE TOPO	TH: : am	243 ino	amin acid	o ac	ids					
50			(xi) SE		ICE D					ID N	0: 3	22:			
		g Il 1	e Th	r As		n Pro	o Gl	u Gl	у Гу	s Tr		u Gl	y Ar	g Thi	r Ala	a Arg
55	Gl	y Se	т Ту	r Gl 2		r Il	e Ly	s Th	r Th 2		a Va	1 G1	u Il	e Xaa		r Asp
	Se	r Le		s Le 5	u Ly	s Ly	s As		r Le 0	u Gl	y Al	a Pr	o Se 4		g Pr	o Ile
60	G1	u As	sp As	p Gl	n Gl	u Va	1 ту	r As	p As	p Va	1 A1	a Gl	u Gl	n As	p As	p Ile

		50					55					60				
5	Ser 65	Ser	His	Ser	Gln	Ser 70	Gly	Ser	Gly	Gly	Ile 75	Phe	Pro	Pro	Pro	Pro 80
	Asp	Asp	Asp	Ile	Tyr 85	Asp	Gly	Ile	Glu	Glu 90	Glu	Asp	Ala	Asp	Asp 95	Gly
10	Phe	Pro	Ala	Pro 100	Pro	Lys	Gln	Leu	Asp 105	Met	Gly	Asp	Glu	Val 110	Tyr	Asp
	Asp	Val	Asp 115	Thr	Ser	Asp	Phe	Pro 120	Val	Ser	Ser	Ala	Glu 125	Met	Ser	Gln
15	Gly	Thr 130	Asn	Val	Gly	Lys	Ala 135	Lys	Thr	Glu	Glu	Lys 140	Asp	Leu	Lys	Lys
20	Leu 145	Lys	Lys	Gln	Xaa	Lys 150	Glu	Xaa	Lys	Asp	Phe 155	Arg	Lys	Lys	Phe	Lys 160
	Tyr	Asp	Gly	Glu	Ile 165	Arg	Val	Leu	Tyr	Ser 170	Thr	Lys	Val	Thr	Thr 175	Ser
25	Ile	Thr	Ser	Lys 180	Lys	Trp	Gly	Thr	Arg 185	Asp	Leu	Gln	Val	Lys 190	Pro	Gly
	Glu	Ser	Leu 195	Glu	Val	Ile	Gln	Thr 200	Thr	Asp	Asp	Thr	Lys 205	Val	Leu	Cys
30	Arg	Asn 210	Glu	Glu	Gly	Lys	Tyr 215	Gly	Tyr	Val	Leu	Arg 220	Ser	Tyr	Leu	Ala
35	Asp 225	Asn	Asp	Gly	Glu	Ile 230	Tyr	Asp	Asp	Ile	Ala 235	Asp	Gly	Cys	Ile	Tyr 240
	Asp	Asn	Asp													
40	(2)							NO: 3 ERIS		:						
4.5				(B) T	YPE:	ami	06 a no a lin	cid	aci	ds					
45								PTIO								
50	1				5					10					Gly 15	
				20					25					30	Gly	
55	Arg	His	Ala 35	Gly	Gly	Gly	Val	His 40	Ile	Glu	Pro	Arg	Tyr 45	Arg	Gln	Phe
		50					55					60			Ser	
50	Leu	Met	Trp	Phe	Trp	Ile	Leu	Trp	Arg	Phe	Trp	His	Asp	Ser	Glu	Glu

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65 70 75 80

Val Leu Gly His Phe Pro Tyr Pro Asp Pro Ser Gln Trp Thr Asp Glu 85 90 95

5
Glu Leu Gly Ile Pro Pro Asp Asp Glu Asp
100
105

Applicant's or agent's file reference number	2004PCT	International application	Unassigned

INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

(PCT Rule 13bis)

A. The indications made below relate to the microorganism referred to in the description on page 73 . line N/A					
B. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet				
Name of depositary institution American Type Culture Col	llection				
Address of depositary institution uncluding postal code and count 10801 University Boulevard Manassas, Virginia 20110-2209 United States of America	(עיט				
Date of deposit March 7, 1997	Accession Number 97923				
C. ADDITIONAL INDICATIONS (leave blank if not applicate	This information is continued on an additional sheet				
	NS ARE MADE (if the indications are not for all designated States)				
E. SEPARATE FURNISHING OF INDICATIONS (leave					
	Bureau later (specify the general nature of the indications, e.g., "Accession				
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Applicant's of agent's the 20041 C.	
reference number	
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INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

(PCT Rule 13bis)

A. The indications made below relate to the microorganism ret on page 73 . line N	ferred to in the description N/A
B. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet
Name of depositary institution American Type Culture (Collection
Address of depositary institution (including postal code and co	ountry)
10801 University Boulevard Manassas. Virginia 20110-2209 United States of America	
Date of deposit May 22, 1997	Accession Number 209071
C. ADDITIONAL INDICATIONS (leave blank if not app.	olicable) This information is continued on an additional sheet
D. DESIGNATED STATES FOR WHICH INDICAT	TIONS ARE MADE (if the indications are not for all designated States)
E. SEPARATE FURNISHING OF INDICATIONS	
The indications listed below will be submitted to the International Number of Deposit's	ional Bureau later (specify the general nature of the indications, e.g., "Accessi
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INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

(PCT Rule 13bis)

A. The indications made below relate to the microorganism refer on page 73 . line N/.	
B. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet :
Name of depositary institution American Type Culture Co	ollection
Address of depositary institution (including postal code and coun	ntry)
10801 University Boulevard Manassas. Virginia 20110-2209 United States of America	
Date of deposit February 25, 1998	Accession Number 209641
C. ADDITIONAL INDICATIONS (leave blank if not application)	able) This information is continued on an additional sheet
D. DESIGNATED STATES FOR WHICH INDICATION	ONS ARE MADE (if the indications are not for all designated States)
	_
E. SEPARATE FURNISHING OF INDICATIONS (leave	
The indications listed below will be submitted to the Internationa Number of Deposit's	Bureau later (specify the general nature of the indications, e.g., "Accession
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Applicant's or agent's file reference number	Z004PCT	International application Unassigned

INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

(PCT Rule 13bis)

. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet
Name of depositary institution American Type	Culture Collection
Address of depositary institution (including postal conference of depositary institution (including postal conference of the conference of	ode and country)
Date of deposit July 24, 1997	Accession Number 209179
C. ADDITIONAL INDICATIONS (leave blan.	k if not applicable) This information is continued on an additional sheet
D. DESIGNATED STATES FOR WHICH I	INDICATIONS ARE MADE (if the indications are not for all designated States)
E. SEPARATE FURNISHING OF INDICA	
E. SEPARATE FURNISHING OF INDICA The indications listed below will be submitted to th	ATIONS (leave blank if not applicable) The international Bureau later (specify the general nature of the indications, e.g., "Accession of the indications of the indication of the indications of the indication of the

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INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

(PCT Rule 13bis)

	A. The indications made below relate to the microorganism referred to in the description on page 77 line N/A			
B. IDENTIFICA	ATION OF DEPOSIT	Further deposits are identified on an additional sheet		
Name of depositary	v institution	and a destroid since		
•		Type Culture Collection		
				
	tary institution (including pos	stal code and country)		
10801 University Manassas, Virgin	nia 20110-2209			
United States of	America			
Date of deposit	March 7, 1997	Accession Number 97924		
	·	77924		
C. ADDITION.	AL INDICATIONS (leave	blank if not applicable) This information is continued on an additional sheet		
		L-)		
D. DESIGNATI	ED STATES FOR WHIC	CH INDICATIONS ARE MADE (if the indications are not for all designated States		
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C. SEPARATE	FURNISHING OF INDI	CATIONS (leave blank if not applicable)		
C. SEPARATE	FURNISHING OF INDI	CATIONS (leave blank if not applicable)		
C. SEPARATE	FURNISHING OF INDI	CATIONS (leave blank if not applicable)		
C. SEPARATE	FURNISHING OF INDI	CATIONS (leave blank if not applicable)		
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C. SEPARATE	FURNISHING OF INDI	CATIONS (leave blank if not applicable)		
E. SEPARATE The indications list Number of Deposit")	FURNISHING OF INDI	CATIONS (leave blank if not applicable) o the International Bureau later (specify the general nature of the indications, e.g., "Access."		
E. SEPARATE The indications list Number of Deposit')	FURNISHING OF INDI- ted below will be submitted to	CATIONS (leave blank if not applicable) o the International Bureau later (specify the general nature of the indications, e.g., "Access The international Bureau later (specify the general nature of the indications, e.g., "Access the indications of the indication of the indications of the indication of		
E. SEPARATE The indications list Number of Deposit')	FURNISHING OF INDI- ted below will be submitted to	CATIONS (leave blank if not applicable) o the International Bureau later (specify the general nature of the indications, e.g., "Access," For International Bureau use only		
E. SEPARATE The indications list Number of Deposit")	FURNISHING OF INDI- ted below will be submitted to	CATIONS (leave blank if not applicable) to the International Bureau later (specify the general nature of the indications, e.g., "Accession to the International Bureau later (specify the general nature of the indications, e.g., "Accession to the International Bureau use only ——For International Bureau use only		
E. SEPARATE The indications list Number of Deposit")	FURNISHING OF INDI- ted below will be submitted to	CATIONS (leave blank if not applicable) the International Bureau later (specify the general nature of the indications, e.g., "Accession of the International Bureau use only — For International Bureau use only — application This sheet was received by the International Bureau on:		
E. SEPARATE The indications list Number of Deposit*) F This sheet wa	FURNISHING OF INDI- ted below will be submitted to	CATIONS (leave blank if not applicable) o the International Bureau later (specify the general nature of the indications, e.g., "Access This sheet was received by the International Bureau on: Authorized officer		

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INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

(PCT Rule 13bis)

B. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet
Name of depositary institution American Type Cultu	
Address of depositary institution (including postal code and 10801 University Boulevard Manassas, Virginia 20110-2209 United States of America	nd country)
Date of deposit March 13, 1997	Accession Number 97958
C. ADDITIONAL INDICATIONS (leave blank if not	t applicable) This information is continued on an additional sheet
D. DESIGNATED STATES FOR WHICH INDIC	CATIONS ARE MADE (if the indications are not for all designated States)
E. SEPARATE FURNISHING OF INDICATION	
E. SEPARATE FURNISHING OF INDICATION The indications listed below will be submitted to the Inter	NS (leave blank if not applicable) mational Bureau later (specify the general nature of the indications, e.g., Accession of the indications of the indication of the indications of the indications of the indication of the

Applicant's or agent's file Z004PCT reference number	International application Unassigned
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INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

(PCT Rule 13bis)

A. The indications made below relate to the microorganism referred to in the description on page 80 . line N/A			
B. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet [7]		
Name of depositary institution American Type Cultur			
Address of depositary institution (including postal code and	country)		
10801 University Boulevard Manassas, Virginia 20110-2209 United States of America			
Date of deposit May 22, 1997	Accession Number 209072		
C. ADDITIONAL INDICATIONS (leave blank if not ap	oplicable) This information is continued on an additional sheet		
D. DESIGNATED STATES FOR WHICH INDICA	TIONS ARE MADE (if the indications are not for all designated States)		
E. SEPARATE FURNISHING OF INDICATIONS	(leave blank if not applicable)		
The indications listed below will be submitted to the International Number of Deposit')	ional Bureau later (specify the general nature of the indications. e.g "Accession		
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INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

(PCT Rule 13bis)

A. The indications made below relate to the microorganism referred to in the description on page 80 , line N/A			
B. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet		
Name of depositary institution American Type Culture Co	ollection		
Address of depositary institution (including postal code and counting 10801 University Boulevard Manassas, Virginia 20110-2209 United States of America	ntry)		
Date of deposit September 4, 1997	Accession Number 209235		
D. DESIGNATED STATES FOR WHICH INDICATI	IONS ARE MADE (if the indications are not for all designated States)		
E. SEPARATE FURNISHING OF INDICATIONS (le. The indications listed below will be submitted to the Internation Number of Deposit')	nal Bureau later (specify the general nature of the indications, e.g., "Accession		
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INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

(PCT Rule 13bis)

A. The indications made below relate to the microorganism on page 84 . line	referred to in the description N/A
B. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet
Name of depositary institution American Type Cultur	re Collection
Address of depositary institution (including postal code and 10801 University Boulevard Manassas, Virginia 20110-2209 United States of America	country)
Date of deposit August 28, 1997	Accession Number 209226
C. ADDITIONAL INDICATIONS (leave blank if not ag	pplicable) This information is continued on an additional sheet
DESIGNATED STATES FOR WHICH INDICA	ATIONS ARE MADE (if the indications are not for all designated States)
E. SEPARATE FURNISHING OF INDICATIONS	
	tional Bureau later (specify the general nature of the indications, e.g., "Accessio
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reference number	

INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

(PCT Rule 13bis)

A. The indications made below relate to the microorganism reference on page 84 line N/	
B. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet
Name of depositary institution American Type Culture C	Collection
Address of depositary institution (including postal code and coulons 10801 University Boulevard Manassas, Virginia 20110-2209 United States of America	intry)
Date of deposit March 13, 1997	Accession Number 97957
C. ADDITIONAL INDICATIONS (leave blank if not applied) D. DESIGNATED STATES FOR WHICH INDICATION	This information is continued on an additional sheet
E. SEPARATE FURNISHING OF INDICATIONS (le	eave blank if not applicable)
The indications listed below will be submitted to the Internation Number of Deposit")	nal Bureau later (specify the general nature of the indications, e.g., "Accession
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INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

(PCT Rule 13bis)

	A. The indications made below relate to the microorganism referred to in the description on page 84 . line N/A								
3. IDENTIFIC	CATION OF DEPOSIT	Further deposits are identified on an additional sheet							
Name of deposita	ry institution								
	American Type	e Culture Collection							
Address of depos	sitary institution (including postal co	Ode and country)							
10801 Universi		,							
	inia 20110-2209								
Office States 0	America								
Date of deposit	May 22, 1997	Accession Number 209073							
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Form PCT/RO/134 (July 1992)

WO 98/42738 PCT/US98/05311

What Is Claimed Is:

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- 1. An isolated nucleic acid molecule comprising a polynucleotide having a nucleotide sequence at least 95% identical to a sequence selected from the group consisting of:
- (a) a polynucleotide fragment of SEQ ID NO:X or a polynucleotide fragment of the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X;
- (b) a polynucleotide encoding a polypeptide fragment of SEQ ID NO:Y or a polypeptide fragment encoded by the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X;
- (c) a polynucleotide encoding a polypeptide domain of SEQ ID NO:Y or a polypeptide domain encoded by the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X;
- (d) a polynucleotide encoding a polypeptide epitope of SEQ ID NO:Y or a polypeptide epitope encoded by the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X;
- (e) a polynucleotide encoding a polypeptide of SEQ ID NO:Y or the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X, having biological activity;
 - (f) a polynucleotide which is a variant of SEQ ID NO:X:
 - (g) a polynucleotide which is an allelic variant of SEQ ID NO:X;
 - (h) a polynucleotide which encodes a species homologue of the SEQ ID NO:Y;
- (i) a polynucleotide capable of hybridizing under stringent conditions to any one of the polynucleotides specified in (a)-(h), wherein said polynucleotide does not hybridize under stringent conditions to a nucleic acid molecule having a nucleotide sequence of only A residues or of only T residues.
- 2. The isolated nucleic acid molecule of claim 1, wherein the polynucleotide fragment comprises a nucleotide sequence encoding a secreted protein.
- 3. The isolated nucleic acid molecule of claim 1, wherein the polynucleotide fragment comprises a nucleotide sequence encoding the sequence identified as SEQ ID NO:Y or the polypeptide encoded by the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X.

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4. The isolated nucleic acid molecule of claim 1, wherein the polynucleotide fragment comprises the entire nucleotide sequence of SEQ ID NO:X or the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X.

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5. The isolated nucleic acid molecule of claim 2, wherein the nucleotide sequence comprises sequential nucleotide deletions from either the C-terminus or the N-terminus.

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- 6. The isolated nucleic acid molecule of claim 3, wherein the nucleotide sequence comprises sequential nucleotide deletions from either the C-terminus or the N-terminus.
- 7. A recombinant vector comprising the isolated nucleic acid molecule of claim 1.
 - 8. A method of making a recombinant host cell comprising the isolated nucleic acid molecule of claim 1.

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- 9. A recombinant host cell produced by the method of claim 8.
- 10. The recombinant host cell of claim 9 comprising vector sequences.
- 11. An isolated polypeptide comprising an amino acid sequence at least 95% identical to a sequence selected from the group consisting of:
 - (a) a polypeptide fragment of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z;
 - (b) a polypeptide fragment of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z, having biological activity;

- (c) a polypeptide domain of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z;
- (d) a polypeptide epitope of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z;
- (e) a secreted form of SEQ ID NO:Y or the encoded sequence included inATCC Deposit No:Z;
 - (f) a full length protein of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z;

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- (g) a variant of SEQ ID NO:Y;
- (h) an allelic variant of SEQ ID NO:Y; or
- (i) a species homologue of the SEQ ID NO:Y.
- 12. The isolated polypeptide of claim 11, wherein the secreted form or the full length protein comprises sequential amino acid deletions from either the C-terminus or the N-terminus.
 - 13. An isolated antibody that binds specifically to the isolated polypeptide of claim 11.
- 14. A recombinant host cell that expresses the isolated polypeptide of claim
 - 15. A method of making an isolated polypeptide comprising:
 - (a) culturing the recombinant host cell of claim 14 under conditions such that said polypeptide is expressed; and
 - (b) recovering said polypeptide.
 - 16. The polypeptide produced by claim 15.
 - 17. A method for preventing, treating, or ameliorating a medical condition, comprising administering to a mammalian subject a therapeutically effective amount of the polypeptide of claim 11 or the polynucleotide of claim 1.
- 25 18. A method of diagnosing a pathological condition or a susceptibility to a pathological condition in a subject comprising:
 - (a) determining the presence or absence of a mutation in the polynucleotide of claim 1; and
 - (b) diagnosing a pathological condition or a susceptibility to a pathological condition based on the presence or absence of said mutation.
 - 19. A method of diagnosing a pathological condition or a susceptibility to a pathological condition in a subject comprising:
 - (a) determining the presence or amount of expression of the polypeptide of claim 11 in a biological sample; and
 - (b) diagnosing a pathological condition or a susceptibility to a pathological condition based on the presence or amount of expression of the polypeptide.

- 20. A method for identifying a binding partner to the polypeptide of claim 11 comprising:
 - (a) contacting the polypeptide of claim 11 with a binding partner; and
- (b) determining whether the binding partner effects an activity of the polypeptide.
 - 21. The gene corresponding to the cDNA sequence of SEQ ID NO:Y.
- 10 22. A method of identifying an activity in a biological assay, wherein the method comprises:
 - (a) expressing SEQ ID NO:X in a cell;
 - (b) isolating the supernatant;
 - (c) detecting an activity in a biological assay; and
 - (d) identifying the protein in the supernatant having the activity.
 - 23. The product produced by the method of claim 22.



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(54) Title: 87 HUMAN SECRETED PROTEINS

(57) Abstract

The present invention relates to 87 novel human secreted proteins and isolated nucleic acids containing the coding regions of the genes encoding such proteins. Also provided are vectors, host cells, antibodies, and recombinant methods for producing human secreted proteins. The invention further relates to diagnostic and therapeutic methods useful for diagnosing and treating disorders related to these novel human secreted proteins.

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87 Human Secreted Proteins

Field of the Invention

This invention relates to newly identified polynucleotides and the polypeptides encoded by these polynucleotides, uses of such polynucleotides and polypeptides, and their production.

Background of the Invention

Unlike bacterium, which exist as a single compartment surrounded by a membrane, human cells and other eucaryotes are subdivided by membranes into many functionally distinct compartments. Each membrane-bounded compartment, or organelle, contains different proteins essential for the function of the organelle. The cell uses "sorting signals," which are amino acid motifs located within the protein, to target proteins to particular cellular organelles.

One type of sorting signal, called a signal sequence, a signal peptide, or a leader sequence, directs a class of proteins to an organelle called the endoplasmic reticulum (ER). The ER separates the membrane-bounded proteins from all other types of proteins. Once localized to the ER, both groups of proteins can be further directed to another organelle called the Golgi apparatus. Here, the Golgi distributes the proteins to vesicles, including secretory vesicles, the cell membrane, lysosomes, and the other organelles.

Proteins targeted to the ER by a signal sequence can be released into the extracellular space as a secreted protein. For example, vesicles containing secreted proteins can fuse with the cell membrane and release their contents into the extracellular space - a process called exocytosis. Exocytosis can occur constitutively or after receipt of a triggering signal. In the latter case, the proteins are stored in secretory vesicles (or secretory granules) until exocytosis is triggered. Similarly, proteins residing on the cell membrane can also be secreted into the extracellular space by proteolytic cleavage of a "linker" holding the protein to the membrane.

Despite the great progress made in recent years, only a small number of genes encoding human secreted proteins have been identified. These secreted proteins include the commercially valuable human insulin, interferon, Factor VIII, human growth hormone, tissue plasminogen activator, and erythropoeitin. Thus, in light of the pervasive role of secreted proteins in human physiology, a need exists for identifying and characterizing novel human secreted proteins and the genes that encode them. This knowledge will allow one to detect, to treat, and to prevent medical disorders by using secreted proteins or the genes that encode them.

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Summary of the Invention

The present invention relates to novel polynucleotides and the encoded polypeptides. Moreover, the present invention relates to vectors, host cells, antibodies, and recombinant methods for producing the polypeptides and polynucleotides. Also provided are diagnostic methods for detecting disorders related to the polypeptides, and therapeutic methods for treating such disorders. The invention further relates to screening methods for identifying binding partners of the polypeptides.

Detailed Description

Definitions

The following definitions are provided to facilitate understanding of certain terms used throughout this specification.

In the present invention, "isolated" refers to material removed from its original environment (e.g., the natural environment if it is naturally occurring), and thus is altered "by the hand of man" from its natural state. For example, an isolated polynucleotide could be part of a vector or a composition of matter, or could be contained within a cell, and still be "isolated" because that vector, composition of matter, or particular cell is not the original environment of the polynucleotide.

In the present invention, a "secreted" protein refers to those proteins capable of being directed to the ER, secretory vesicles, or the extracellular space as a result of a signal sequence, as well as those proteins released into the extracellular space without necessarily containing a signal sequence. If the secreted protein is released into the extracellular space, the secreted protein can undergo extracellular processing to produce a "mature" protein. Release into the extracellular space can occur by many mechanisms, including exocytosis and proteolytic cleavage.

As used herein, a "polynucleotide" refers to a molecule having a nucleic acid sequence contained in SEQ ID NO:X or the cDNA contained within the clone deposited with the ATCC. For example, the polynucleotide can contain the nucleotide sequence of the full length cDNA sequence, including the 5' and 3' untranslated sequences, the coding region, with or without the signal sequence, the secreted protein coding region, as well as fragments, epitopes, domains, and variants of the nucleic acid sequence. Moreover, as used herein, a "polypeptide" refers to a molecule having the translated amino acid sequence generated from the polynucleotide as broadly defined.

In the present invention, the full length sequence identified as SEQ ID NO:X was often generated by overlapping sequences contained in multiple clones (contig

analysis). A representative clone containing all or most of the sequence for SEQ ID NO:X was deposited with the American Type Culture Collection ("ATCC"). As shown in Table 1, each clone is identified by a cDNA Clone ID (Identifier) and the ATCC Deposit Number. The ATCC is located at 10801 University Boulevard, Manassas, Virginia 20110-2209, USA. The ATCC deposit was made pursuant to the terms of the Budapest Treaty on the international recognition of the deposit of microorganisms for purposes of patent procedure.

A "polynucleotide" of the present invention also includes those polynucleotides capable of hybridizing, under stringent hybridization conditions, to sequences contained in SEQ ID NO:X, the complement thereof, or the cDNA within the clone deposited with the ATCC. "Stringent hybridization conditions" refers to an overnight incubation at 42° C in a solution comprising 50% formamide, 5x SSC (750 mM NaCl, 75 mM sodium citrate), 50 mM sodium phosphate (pH 7.6), 5x Denhardt's solution, 10% dextran sulfate, and 20 μ g/ml denatured, sheared salmon sperm DNA, followed by washing the filters in 0.1x SSC at about 65°C.

Also contemplated are nucleic acid molecules that hybridize to the polynucleotides of the present invention at lower stringency hybridization conditions. Changes in the stringency of hybridization and signal detection are primarily accomplished through the manipulation of formamide concentration (lower percentages of formamide result in lowered stringency); salt conditions, or temperature. For example, lower stringency conditions include an overnight incubation at 37°C in a solution comprising 6X SSPE (20X SSPE = 3M NaCl; 0.2M NaH₂PO₄; 0.02M EDTA, pH 7.4), 0.5% SDS, 30% formamide, 100 ug/ml salmon sperm blocking DNA; followed by washes at 50°C with 1XSSPE, 0.1% SDS. In addition, to achieve even lower stringency, washes performed following stringent hybridization can be done at higher salt concentrations (e.g. 5X SSC).

Note that variations in the above conditions may be accomplished through the inclusion and/or substitution of alternate blocking reagents used to suppress background in hybridization experiments. Typical blocking reagents include Denhardt's reagent, BLOTTO, heparin, denatured salmon sperm DNA, and commercially available proprietary formulations. The inclusion of specific blocking reagents may require modification of the hybridization conditions described above, due to problems with compatibility.

Of course, a polynucleotide which hybridizes only to polyA+ sequences (such as any 3' terminal polyA+ tract of a cDNA shown in the sequence listing), or to a

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complementary stretch of T (or U) residues, would not be included in the definition of "polynucleotide," since such a polynucleotide would hybridize to any nucleic acid molecule containing a poly (A) stretch or the complement thereof (e.g., practically any double-stranded cDNA clone).

The polynucleotide of the present invention can be composed of any polyribonucleotide or polydeoxribonucleotide, which may be unmodified RNA or DNA or modified RNA or DNA. For example, polynucleotides can be composed of single-and double-stranded DNA, DNA that is a mixture of single- and double-stranded regions, single- and double-stranded RNA, and RNA that is mixture of single- and double-stranded regions, hybrid molecules comprising DNA and RNA that may be single-stranded or, more typically, double-stranded or a mixture of single- and double-stranded regions. In addition, the polynucleotide can be composed of triple-stranded regions comprising RNA or DNA or both RNA and DNA. A polynucleotide may also contain one or more modified bases or DNA or RNA backbones modified for stability or for other reasons. "Modified" bases include, for example, tritylated bases and unusual bases such as inosine. A variety of modifications can be made to DNA and RNA; thus, "polynucleotide" embraces chemically, enzymatically, or metabolically modified forms.

The polypeptide of the present invention can be composed of amino acids joined to each other by peptide bonds or modified peptide bonds, i.e., peptide isosteres, and may contain amino acids other than the 20 gene-encoded amino acids. The polypeptides may be modified by either natural processes, such as posttranslational processing, or by chemical modification techniques which are well known in the art. Such modifications are well described in basic texts and in more detailed monographs, as well as in a voluminous research literature. Modifications can occur anywhere in a polypeptide, including the peptide backbone, the amino acid side-chains and the amino or carboxyl termini. It will be appreciated that the same type of modification may be present in the same or varying degrees at several sites in a given polypeptide. Also, a given polypeptide may contain many types of modifications. Polypeptides may be branched, for example, as a result of ubiquitination, and they may be cyclic, with or without branching. Cyclic, branched, and branched cyclic polypeptides may result from posttranslation natural processes or may be made by synthetic methods. Modifications include acetylation, acylation, ADP-ribosylation, amidation, covalent attachment of flavin, covalent attachment of a heme moiety, covalent attachment of a nucleotide or nucleotide derivative, covalent attachment of a lipid or lipid derivative, covalent attachment of phosphotidylinositol, cross-linking, cyclization, disulfide bond formation, demethylation, formation of covalent cross-links, formation of cysteine,

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formation of pyroglutamate, formylation, gamma-carboxylation, glycosylation, GPI anchor formation, hydroxylation, iodination, methylation, myristoylation, oxidation, pegylation, proteolytic processing, phosphorylation, prenylation, racemization, selenoylation, sulfation, transfer-RNA mediated addition of amino acids to proteins such as arginylation, and ubiquitination. (See, for instance, PROTEINS - STRUCTURE AND MOLECULAR PROPERTIES, 2nd Ed., T. E. Creighton, W. H. Freeman and Company, New York (1993); POSTTRANSLATIONAL COVALENT MODIFICATION OF PROTEINS, B. C. Johnson, Ed., Academic Press, New York, pgs. 1-12 (1983); Seifter et al., Meth Enzymol 182:626-646 (1990); Rattan et al., Ann NY Acad Sci 663:48-62 (1992).)

"SEQ ID NO:X" refers to a polynucleotide sequence while "SEQ ID NO:Y" refers to a polypeptide sequence, both sequences identified by an integer specified in Table 1.

"A polypeptide having biological activity" refers to polypeptides exhibiting activity similar, but not necessarily identical to, an activity of a polypeptide of the present invention, including mature forms, as measured in a particular biological assay, with or without dose dependency. In the case where dose dependency does exist, it need not be identical to that of the polypeptide, but rather substantially similar to the dose-dependence in a given activity as compared to the polypeptide of the present invention (i.e., the candidate polypeptide will exhibit greater activity or not more than about 25-fold less and, preferably, not more than about tenfold less activity, and most preferably, not more than about three-fold less activity relative to the polypeptide of the present invention.)

25 Polynucleotides and Polypeptides of the Invention

FEATURES OF PROTEIN ENCODED BY GENE NO: 1

The translation product of this gene shares sequence homology with nucleolin, which is thought to be important in macromolecule binding, as well as some membrane proteins. Preferred polypeptide fragments comprise the amino acid sequence:

DPEAADSGEPQNKRTPDLPEEEYVKEEIQENEEAVKKMLVEATREFEEVVVDES (SEQ ID NO:239); QKLKRKAEEDPEAADSGEPQNKRTPDLPEEEYVKEEIQENEE AVKKMLVEATREFEEVVVDES (SEQ ID NO:240); KAMEKSSLTQHSWQSLKDR YLKHLRGQEHKYLLGDAPVSPSSQKLKRKAEEDPEAADSGEPQNKRTPDLPEE

35 EYVKEEIQENEEAVKKMLVEATREFEEVVVDESPPDFEIHI (SEQ ID NO:241). Also preferred are the polynucleotide fragments encoding these polypeptide fragments.

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This gene maps to chromosome 16, and therefore can be used as a marker in linkage analysis for chromosome 16.

This gene is expressed primarily in brain and kidney and to a lesser extent in wide range of tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cell-cell interaction or cell-matrix interaction. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the brain and kidney, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., brain and other tissue of the nervous system, and kidney, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:125 as residues: Met-1 to Trp-10.

The tissue distribution and homology to nucleolin indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment/diagnosis of diseases involving cell-cell interaction or cell-extracellular matrix interaction.

FEATURES OF PROTEIN ENCODED BY GENE NO: 2

The translation product of this gene shares sequence homology with a porcine zona pellucida protein ZPDS.1711. (See Accession No. R39356.) These two proteins have weak homology with *Drosophila* commissureless and metal homeostasis proteins which are thought to be important in controlling growth cone guidance across the CNS midline and protecting cells against reactive oxygen toxicity, thus, based on homology, it is likely that this gene also be involved in development. Preferred polypeptide fragments comprise the amino acid sequence: LPSYDEAERTKAEATIPLVPGRDEDF VGRDDFDDADQLRIGNDGIFMLTFFMAFLFNWIGFFLSFCLTTSAAGRYGAISG FGLSLIKWILIVRFSTYFPGYFDGQYWLWWVFLVLGFLLFLRGFINYAKVRKM PETFSNLPRTRVLFI (SEQ ID NO:242); and/or AGRYGAISGFGLSLIKWILIVRFS (SEQ ID NO:243). Also preferred are polynucleotide fragments encoding these polypeptide fragments. This gene maps to chromosome 5, and therefore can be used in linkage analysis as a marker for chromosome 5.

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This gene is expressed primarily in kidney, adrenal gland, brain and to a lesser extent in wide range of tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, fertilization control or tissue damages by metabolites or other toxic agents. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the reproductive and urosecretion system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., kidney, adrenal gland, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to zona pellucida protein indicates that polynucleotides and polypeptides corresponding to this gene are useful for fertility control such as controceptive development. The homology with metal homeostasis and commissureless genes indicates the gene's function in spermatozoa guidance and protection. It would also be useful for the treatment/diagnosis of tissue damages caused by toxic metabolites and other agents since the gene product is also expressed in urosecretive tissues.

25 FEATURES OF PROTEIN ENCODED BY GENE NO: 3

This gene is expressed primarily in liver and to a lesser extent in placenta. Preferred polypeptide fragments comprise the amino acid sequence: MKHLSAWNFT KLTFLQLWEI FEGSVENCQTLTSYSKLQIKYTFSRGSTFYI (SEQ ID NO:244). Also preferred are polynucleotide fragments encoding these polypeptide fragments.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, digestive and nutrient transport/utilization disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the digestive and

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circulatory system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., liver, and placenta, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution in liver and placenta indicates that the protein product is either an extracellular enzyme or a molecule carrier. Therefore, polynucleotides and polypeptides corresponding to this gene are useful for diagnosis/treatment of digestive and nutrient transport/utilization disorders, including malabsorption and malnutrition.

FEATURES OF PROTEIN ENCODED BY GENE NO: 4

This gene shares homology with the sap47 gene of Drosophila melanogaster, a gene which codes for a conserved neuronal protein associated with synaptic terminals. (See 15 Mol. Brain Res. 32:45-54 (1995); see also, Accession No. 929571.) Thus, based on homology, the gene of the present invention also should be associated with synaptic terminals. Preferred polypeptide fragments comprise the amino acid sequence: FSSDFRTSPWESRRVESKATSARCGLWGSGPRRRPASGMFRGLSSWLGLQQPVAGGGQPNGDAPPEQPSETVAESAEEELQQAGDQELLHQAKDFGNYLFNFASA 20 ATKKITESVAETAQTIKKSVEEGKIDGIIDKTIIGDFOKEOKKFVEEOHTKKSEA AVPPWVDTNDEETIQQQILALSADKRNFLRDPPAGVQFNFDFDQMYPVALVML (SEQ ID NO:245); MRFALVPKLVKEEVFWRNYFYRVSLIKQSAQLTALAAQQQA AGKGGEEQ (SEQ ID NO:246); STSPGVSEFVSDAFDACNLNQEDLRKEMEQL VLDKKQEETAVLEEDSADWEKELQQELQEYEVVTESEKRDENWDK (SEQ ID 25 NO:247); SPWESRRVESKATSARCGLWGSGPRRRPASGMFRGLSSWLGLQQ PVAGGGQPNGDAPPEQPS (SEQ ID NO:248); PVAGGGQPNGDAPPEQPSETV ESAEEELQQAGDQELLHQAKDFGNYLFNFASAATKKITESVAE (SEQ ID NO: 249); and/or FQKEQKKFVEEQHTKKSEAAVPPWVDTNDEETIQQQILALSADKR NFLRDPPAGVQFNFDFDQMYPVALVML (SEQ ID NO:250). Also preferred are polynucleotide fragments encoding these polypeptide fragments. 30

This gene is expressed primarily in kidney pyramids and to a lesser extent in lung and other tissues of various types. This gene fluxes calcium in human aortic smooth muscle cells, and therefore is involved in signal transduction.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are

not limited to, renal and nervous disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the kidney and/or nervous system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., kidney, lung, brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution in kidney and lung and homology with sap47 indicates that the protein product has regulatory or direct functions in molecular exchange with body fluids and nervous system signaling. Polynucleotides and polypeptides corresponding to this gene are useful for treatment of disorders in kidney and nervous system.

FEATURES OF PROTEIN ENCODED BY GENE NO: 5

The translation product of this gene shares sequence homology with the mouse Ly-9.2 antigen which is thought to be an important cell surface marker in lymphoids, myeloids and hematopoietic progenitors. (See Accession No. gil198932.) Preferred polypeptide fragments comprise the amino acid sequence: PFICVARNPVSRNFSSPI LARKLCEGAA (SEQ ID NO:251); and/or KEDPANTVYSTVEIPKKMENPHSLLT MPDTPRL (SEQ ID NO:252). Also preferred are polynucleotide fragments encoding these polypeptide fragments. Based on homology, it is likely that this gene is also a cell surface marker, involved in hematopoiesis.

This gene is expressed primarily in activated macrophages, monocytes and T-cells and to a lesser extent in spleen and bone marrow.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune and hematopoietic disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and hematopoietic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., blood cells, and bone marrow, and cancerous and wounded

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tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:129 as residues: Lys-26 to Tyr-33, Arg-44 to Ile-49, Ser-53 to Lys-71, Lys-86 to Pro-91.

The tissue distribution and homology to Ly-9.2 surface immunoglobulin family indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis of immune and hematopoietic disorders. Polypeptides and polynucleotides corresponding to this gene are also be used as a marker for leukemia or a modulator of the functions of the cells of macrophage/monocyte or T-cell types.

FEATURES OF PROTEIN ENCODED BY GENE NO: 6

The translation product of this gene shares sequence homology with the 15 Drosophila glutactin gene which is thought to be important in cell-cell interaction or cell-extracellular matrix contact.

This gene is expressed primarily in colon tissue, aorta endothelial cells and to a lesser extent in skin, breast tissue and T-cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of these tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, diseases of the gastrointestinal tract, vascular system or T-cell development. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the digestive system, cardiovascular system, and immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., colon, cardiovascular tissue, skin, mammary tissue, and blood cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to glutactin indicates that polynucleotides and polypeptides corresponding to this gene are useful for the development and maintenance of the integrity of the basal membrane in the gastrointestinal tract and

cardiovascular system. The expression in T-cells also indicate the protein may be involved in T-cell adhesion, cell-cell interaction and development.

FEATURES OF PROTEIN ENCODED BY GENE NO: 7

The translation product of this gene shares sequence homology with MURF4 protein, an ATPase homolog, which is thought to be important in ATP hydrolysis.

This gene is expressed primarily in breast tissue.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, breast cancer and non-neoplastic breast diseases. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the breast tissue, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., mammary tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to MURF4 gene indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of neoplastic or non-neoplastic breast diseases because ATPase like protein may be involved in changed metabolic states of the breast.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 8

This gene shares homology to the alcohol dehydrogenase gene. Preferred polypeptide fragments comprise comprise the amino acid sequence: ASAVLLDLPNSG GEAQAKKLGNNCVFAPADVTSEKDVQTALALAKGKFGRVDVAVNCAGIAVAS KTYNLKKGQTHTLEDFQRVLDVNLMGTFNVIRLVAGEMGQNEPDQGGQRGVI INTASVAAFEGQVGQAAYSASKGGIVGMTLPIARDLAPIGIRVMTIAPGLFGTPL LTSLPEKVCNFLASQVPFPSRLGDPAEYAHLVQAIIENPFLNGEVIRLDGAIRMQ P (SEQ ID NO:253); and/or SVAAFEGQVGQAAYSASKGGIVGMTLPIA (SEQ ID NO:254). Polynucleotides encoding these fragements are also encompassed by the invention. Other groups have also recently cloned this gene, recognizing its homology to alcohol dehydrogenase. (See Accession No. 1778355.) Moreover, a second group

recently cloned the mouse homologue of this gene. (See Accession No. 2078284.) They found that the mouse homologue binds to amyloid beta-peptide and mediates neurotoxicity in Alzheimer's disease, calling the protein ERAB. This gene maps to chromosome X, and therefore can be used in linkage analysis as a marker for chromosome X. Therefore, mutations in the translated product of this gene may be involved in Alzheimer's disease in humans, as well as other sex linked diseases. This gene can be used as a diagnostic marker for these diseases.

Preferred epitopes include those comprising a sequence shown in SEQ ID NO:132 as residues: Arg-45 to Ser-53.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 9

The translation product of this gene shares week sequence homology with rat N-methyl-D-aspartate receptor subunit and other proline-rich proteins which are thought to be important in neurotransmission or protein-protein intereaction.

This gene is expressed primarily in synovial hypoxia and to a lesser extent in ovary, senescent cells and brain.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, synovial hypoxia. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the synovia and brain, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., synovial tissue, ovary and other reproductive tissue, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution in synovial hypoxia and nerve tissues, and homology to N-methyl-D-aspartate receptor subunit and other proline-rich proteins indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and intervention of synovial hypoxia and other synovial disorders.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 10

This gene is expressed primarily in prostate and to a lesser extent in placenta and ovary.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, male and female infertility, cancer, and other hyperproliferative disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, 10 particularly of the reproductive system and neoplasia, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., prostate, placenta, ovary and other reproductive tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to 15 the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:134 as residues: Pro-17 to Met-23, Ala-30 to Trp-38, Ile-49 to Trp-54, Lys-68 to Gly-74, Thr-93 to Gly-99, Met-126 to Glu-132, Gly-173 to Ser-178, Lys-205 to Tyr-214. 20

The tissue distribution of this gene in the prostate, placenta and ovary indicates that this gene product is useful for treatment/diagnosis of male or female infertility, endocrine disorders, fetal deficiencies, ovarian failure, amenorrhea, ovarian cancer, benign prostate hyperplasia, prostate cancer, and other forms of cancer of the reproductive system.

FEATURES OF PROTEIN ENCODED BY GENE NO: 11

This gene is expressed primarily in the thyroid and to a lesser extent in the pineal gland. This gene maps to chromosome 10, and therefore can be used as a marker in linkage analysis for chromosome 10. Preferred polypeptide fragments comprise the amino acid sequence: HPIEWAINAATLSQFY (SEQ ID NO:256); CWIKYCLTLMQN AQLSMQDNIG (SEQ ID NO:257); KVSYLRPLDFEEARELFLLGQHYVF (SEQ ID NO:258); MERRCKMHKRXIAMLEPLTVDLNPQ (SEQ ID NO:259); and/or SHIV KKINNLNKSALKY YQLFLD (SEQ ID NO:260). Also preferred are polynucleotides encoding these polypeptide fragments.

Therefore, polynucleotides and polypeptides of the invention are useful as

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reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune, thyroid and pineal gland disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and endocrine systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., thyroid and pineal gland, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:135 as residues: Ser-2 to Ser-8, Thr-38 to Arg-44.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treating/detecting immune disorders such as arthritis, asthma, immune deficiency diseases (e.g., AIDS), and leukemia, as well as treating/detecting thymus disorders (e.g., Graves Disease, lymphocytic thyroiditis, hyperthyroidism, and hypothyroidism), and treating/detecting pineal gland disorders (e.g., circadian rhythm disturbances associated with shift work, jet lag, blindness, insomnia and old age).

FEATURES OF PROTEIN ENCODED BY GENE NO: 12

This gene is expressed primarily in lung and tonsils.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, pulmonary or immune disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the pulmonary and immune systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., pulmonary tissue, and tonsils, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily

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fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:136 as residues: Glu-28 to Gly-49.

The tissue distribution of this gene only in lung indicates that it could play a role in the treatment/detection of lung lymphoma or sarcoma formation, pulmonary edema and embolism, bronchitis and cystic fibrosis. Its expression in tonsils indicates a potential role in the treatment/detection of immune disorders such as arthritis, asthma, immune deficiency diseases (e.g., AIDS), and leukemia, in addition to the treatment/detection of tonsillitis.

FEATURES OF PROTEIN ENCODED BY GENE NO: 13

This gene is expressed primarily in lymphoid, myeloid and erythroid cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, hematopoietic and immune disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the hematopoietic and immune systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., blood cells, myeloid cells, and bone marrow, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The predominant tissue distribution of this gene in hematopoietic cell types indicates that the gene could be important for the treatment or detection of immune or hematopoietic disorders including arthritis, asthma, immunodeficiency diseases and leukemia. Preferred embodiments of the present invention are polypeptide fragments comprising the amino acid sequence: FTHLSTCLLSLLLVRMSGFLLLARASPSI CALDSSCFVEYCSSYSSSCFLHQHFPSLLDHLCQ (SEQ ID NO:261); or FLLL ARASPSICALDSSCFVQEY (SEQ ID NO:262). Also preferred are polynucleotide fragments encoding these polypeptide fragments.

FEATURES OF PROTEIN ENCODED BY GENE NO: 14

This gene is homologous to the *Drosophila Regena* (Rga) gene. (See Accession No. 1658504.) This *Drosophila* gene is thought to be a homolog of the global negative

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transcriptional regulator NOT2 (CDC36) from yeast, which modifies gene expression and suppresses position effect variegation. Preferred polypeptide fragments comprise the amino acid sequence: PDGRVTNIPQGMVTDQFGMIGLLTFIRAAETDPGMVHL ALGSDLTTLGLNLNS (SEQ ID NO:263); VHLALGSDLTTLGLNLNSPENLYP (SEQ ID NO:265); EDLLFYLYYMNGGDVLQLLAAVELFNRDWRYHKEERVWI TR (SEQ ID NO:264); and/or HNEDFPALPGS (SEQ ID NO:266).

This gene is expressed primarily in placenta and to a lesser extent in infant brain.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, neurodegenerative and developmental disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the neurological system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., placenta, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:138 as residues: Leu-9 to Tyr-15, Asp-34 to Gln-46, Pro-51 to Asp-57, Gly-88 to Thr-104, Thr-123 to Ser-128.

The tissue distribution of this gene indicates that it could be used in the detection and/or treatment of neurological disorders such as such as Alzheimer's Disease, Parkinson's Disease, Huntington's Disease, schizophrenia, mania, dementia, paranoia, obsessive compulsive disorder, and panic disorder.

FEATURES OF PROTEIN ENCODED BY GENE NO: 15

This gene is expressed primarily in adrenal gland tumor and osteoclastoma.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, endocrine and bone disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for

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differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the endocrine system and in bone, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., adrenal gland, and bone, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:139 as residues: Ile-52 to Trp-57.

The tissue distribution of this gene indicates that it may be involved in the treatment and/or detection of adrenal gland tumors, osteosarcomas, endocrine disorders and bone disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 16

The translation product of this gene shares sequence homology with the FK506 binding protein, a protein which plays an important role in immunosupression. (See Accession No. M75099.) Specifically, a 12-kDa FK506-binding protein (FKBP-12) is a cytosolic receptor for the immunosuppressants FK506 and rapamycin. (See, Proc. Natl. Acad. Sci. 88: 6677-6681 (1991).) Thus, based on homology, it is likely that this gene also has immunosuppression activity. Preferred polypeptides comprise the amino acid sequence: GRIIDTSLTRDPLVIELGQKQVIPGLEQSLLDMCVGEKRRAIIPSH LAYGKRGFPPSVPADAVVQYDVELIALIR (SEQ ID NO:267); and/or IHYTGSLV DGR IIDTS (SEQ ID NO:268). Also preferred are the polynucleotide fragments encoding these polypeptides.

This gene is expressed primarily in melanocytes.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cancer and other hyperproliferative disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system and cancer, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., melanocytes, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to

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the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:140 as residues: Ala-118 to Phe-124, Arg-178 to Lys-201.

The tissue distribution and homology to the FK506 binding proteins which are believed to a role in immunosupression mediated by the immunosupressant drugs rapamycin and cyclosporin, indicates that this gene could serve as a novel target for the identification of novel immunosupressant drugs.

10 FEATURES OF PROTEIN ENCODED BY GENE NO: 17

The translation product of this gene shares sequence homology with the rat calcium-activated potassium channel rSK3, which is thought to be important in regulating vascular tone. (See Accession No. gil2564072, gil1575663, and gil1575661.) Although homologous to these proteins, this gene contains an 18 amino acid insert, not previously identified in the homologs. Preferred polypeptide fragments comprise the amino acid sequence: CESPESPAQPSGSSLPAWYH (SEQ ID NO:269). Also preferred are the polynucleotide fragments encoding these polypeptides.

This gene is expressed primarily in B-cells, frontal cortex and endothelial cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cardiovascular (hyper/hypotension, asthma, pulmonary edema, pneumonia, heart disease, restenosis, atherosclerosis, stoke, angina and thrombosis) and neurological disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the cardiovascular and nervous systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., blood cells, brain and other tissue of the nervous system, and endothelium, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:141 as residues: Glu-72 to Gly-82, His-90 to Val-95, Gln-168 to Lys-174, Val-202 to Ser-212.

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The tissue distribution and homology to calcium-activated potassium channels indicates that polynucleotides and polypeptides corresponding to this gene are useful for the diagnosis and treatment of vascular disorders (hyper/hypotension, athesma, pulmonary edema, pneumonia, heart disease, restenosis, atherosclerosis, stoke, angina and thrombosis).

FEATURES OF PROTEIN ENCODED BY GENE NO: 18

This gene is expressed primarily in smooth muscle and to a lesser extent in brain (amygdala, corpus colosum, hippocampus).

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cardiovascular (hypertension, heart disease, athesma, pulmonary edema, restenosis, atherosclerosis, stoke, angina, thrombosis, and wound healing), and neurological disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the cardiovascular and neurological systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., smooth muscle, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:142 as residues: Lys-43 to Arg-49, Tyr-58 to Glu-65.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of cadiovascular disorders (hypertension, heart disease, athesma, pulmonary edema, restenosis, atherosclerosis, stoke, angina, thrombosis, and wound healing). Expression in brain indicates a role in the treatment and diagnosis of behavioral or neurological disorders, such as depression, schizophrenia, Alzheimer's disease, mania, dementia, paranoia, and addictive behavior.

35 FEATURES OF PROTEIN ENCODED BY GENE NO: 19

This gene is expressed primarily in T-cells (Jurkats, resting, activated, and

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anergic T-cells), endothelial cells, pineal gland, and to a lesser extent in a variety of other tissues and cell types. Preferred polypeptide fragments comprise the amino acid sequence: EEAGAGRRCSHGGARPAGLGNEGLGLGGDPDHTDTGSRSKQRINN WKESKHKVIMASASARGNQDKDAHFPPPSKQSLLFCPKSKLHIHRAEISK (SEQ ID NO:270); and/or SKQRINNWKESKHKVIMASASAR (SEQ ID NO:271). Also preferred are the polynucleotide fragments encoding these polypepides.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, inflammation, immune and cardiovascular disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, neurological and vascular systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, endothelial cells, and pineal gland, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:143 as residues: Phe-71 to Arg-76, Pro-82 to His-87, Glu-103 to Ala-111.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the diagnosis and treatment of immune disorders including: leukemias, lymphomas, auto-immune, immuno-supressive (e.g. transplantation) and immunodeficiencies (e.g. AIDS) and hematopoietic disorders. In addition, expression in the pineal gland might suggest a role in the diagnosis of specific brain tumors and treatment of neurological disorders. Endothelial cell expression might suggest a role in cadiovascular or respiratory/pulmonary disorders or infections (athesma, pulmonary edema, pneumonia).

FEATURES OF PROTEIN ENCODED BY GENE NO: 20

This gene is expressed primarily in brain and embryo and to a lesser extent in leukocytes. This gene maps to chromosome 15, and therefore can be used as a marker in linkage analysis to chromosome 15.

Therefore, polynucleotides and polypeptides of the invention are useful as

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reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, developmental and neurological disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the central nervous and immune systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:144 as residues: Met-1 to Gly-8.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of immune disorders including: leukemias, lymphomas, auto-immune, immuno-supressive (e.g. transplantation) and immunodeficiencies (e.g. AIDS) and hematopoietic disorders. The expression in the brain -- and in particular the fetal brain -- would suggest a possible role in the treatment and diagnosis of developmental and neurodegenerative diseases of the brain and nervous system (depression, schizophrenia, Alzheimer's disease, mania, dementia, paranoia, and addictive behavior).

FEATURES OF PROTEIN ENCODED BY GENE NO: 21

This gene is expressed primarily in brain, kidney, lung, liver, spleen, and a variety of leukocytes (especially T-cells) and to a lesser extent in a variety of other tissues and cell types.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, leukemias, lymphomas, autoimmune, immunosuppressive, and immunodeficiencies, hematopoietic disorders, as well as renal disorders, and neoplasms. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the renal, pulmonary, immune, and central nervous systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g.,

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brain and other tissue of the nervous system, kidney, pulmonary tissue, liver, spleen, and blood cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of renal conditions, such as acture renal failure, kidney fibrosis, and kidney tubule regeneration. The expression in leukocytes and other immune tissues indicates a role in immune disorders including: leukemias, lymphomas, auto-immune, immuno-supressive (e.g. transplantation) and immunodeficiencies (e.g. AIDS) and hematopoietic disorders. The expression in the brain -- and in particular the fetal brain -- indicates a possible role in the treatment and diagnosis of developmental and neurodegenerative diseases of the brain and nervous system (depression, schizophrenia, Alzheimer's disease, mania, dementia, paranoia, and addictive behavior).

FEATURES OF PROTEIN ENCODED BY GENE NO: 22

This gene is expressed primarily in skin (fetal epithelium, keratinocytes and skin). This gene also maps to chromosome 19, and therefore can be used in linkage analysis as a marker for chromosome 19.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, skin cancers (e.g., melanomas), eczema, psoriasis or other disorders of the skin. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the skin, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., skin and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:146 as residues: Pro-28 to Glu-35, Ser-39 to Phe-44, Ala-94 to Gln-99.

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The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of skin cancers (e.g., melanomas), eczema, psoriasis or other disorders of the skin.

5 FEATURES OF PROTEIN ENCODED BY GENE NO: 23

This gene maps to chromosome 11. Another group recently isolated this same gene, associating the sequence to the region thought to harbor the gene involved in Multiple Endocrine Neoplasia Type 1, or MEN 1. (See Accession No. 2529721 and Genome Res. 7(7), 725-735 (1997), incorporated herein by reference in its entirety.) Preferred polypeptide fragments comprise the amino acid sequence: LFHWACLNERA AOLPRNTAXAGYQCPSCNGPS (SEQ ID NO:272).

This gene is expressed primarily in epididymus, pineal gland, T-cells, as well as fetal epithelium, lung and kidney.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune, metabolic mediated disorders, and MEN. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, renal, neurological and pulmonary systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., epididymus and other reproductive tissue, pineal gland, T-cells and other blood cells, epithelium, lung, and kidney, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of developmental deficiencies or abnormalities as well as a host of different disorders which arise as a result of conditions in the indicated tissues or cell types. An area of particular interest is in the treatment and diagnosis of immune disorders including: leukemias, lymphomas, auto-immune, immuno-supressive (e.g. transplantation) and immunodeficiencies (e.g. AIDS) and hematopoietic disorders. The expression in the brain, and in particular the fetal brain, would suggest a possible role in the treatment and diagnosis of

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developmental and neurodegenerative diseases of the brain and nervous system (depression, schizophrenia, Alzheimer's disease, mania, dementia, paranoia, and addictive behavior). Respiratory/pulmonary disorders, such as athesma, pulmonary edema are also potential therapeutic areas, as well as renal conditions such as acute renal failure, kidney fibrosis and kidney tubule regeneration. Moreover, this gene can be used in the treatment and/or detection of MEN I.

FEATURES OF PROTEIN ENCODED BY GENE NO: 24

This gene is expressed primarily in fetal spleen.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, leukemia, lymphoma, AIDS, hematopoeitic disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and hematopoietic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., spleen and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of immune disorders including: leukemias, lymphomas, auto-immune, immuno-supressive (e.g. transplantation) and immunodeficiencies (e.g. AIDS) and hematopoietic disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 25

A closely related homolog of this gene was recently cloned by another group,

calling the gene CDO, an oncogene-, serum-, and anchorage-regulated member of the
Ig/fibronectin type III repeat family. (See Accession No. 2406628, and J. Cell Biol.
138(1): 203-213 (1997), herein incorporated by reference in its entirety.) Preferred
polypeptide fragments comprise the amino acid sequence: FYIYYRPTDSDNDSDYKK
DMVEGDKYWHSISHLQPETSYDIKMQCFNEGGESEFSNVMICETKARKSSGQP

GRLPPPTLAPPQPPLPETIERPVGTGAMVARSSDLPYLIVGVVLGSIVLIIVTFIPF
CLWRAWSKQKHTTDLGFPRSALPPSCPYTMVPLGGLPGHQAVDSPTSVASVD

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GPVLM (SEQ ID NO:273); or YIYYRPTDSDNDSDYKKDMVEGDKYWHSISHLQ PETSYDIKMQCFNEGGESEFSNVMICETKARKS (SEQ ID NO:274).

This gene is expressed primarily in fetal lung and kidney, human embryo and osteoclastoma stromal cells and to a lesser extent in a variety of other tissues and cell types.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, developmental disorders and cancers, as well as pulmonary and renal disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the respiratory/pulmonary, skeletal and renal systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., lung, kidney, embryonic tissue, and bone cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:149 as residues: Thr-5 to Pro-18, Ala-76 to Thr-84.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the detection and treatment of: osteoperosis, fracture, osteosarcoma, ossification, and osteonecrosis, as well as respiratory/pulmonary disorders, such as athesma, pulmonary edema, and renal conditions such as acute renal failure, kidney fibrosis and kidney tubule regeneration.

FEATURES OF PROTEIN ENCODED BY GENE NO: 26

This gene is homologous to the HIV envelope glycoprotein. (See Accession No. 2641463.) Preferred polypeptide fragments comprise the amino acid sequence: NVRALLHRMPEPPKINTAKFNNNKRKNLSL (SEQ ID NO:275).

This gene is expressed primarily in pineal gland and skin, and to a lesser extent in lung.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are

not limited to, neurological and behavior disorders; respiratory/pulmonary disorders, such as athesma, pulmonary edema; skin conditions such as eczema, psoriasis, acne and skin cancer, as well as AIDS. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the central nervous and respiratory systems, as well as skin and AIDS, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., blood cells, pineal gland, epidermis, and pulmonary tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:150 as residues: Gln-15 to Gln-20.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of conditions which affect the above tissues, such as: skin cancer, eczema, psoriasis, acne, athesma, pulmonary edema, neuro-degenerative or developmental disorders such as Alzheimer's, depression, schizophrenia, dementia, and AIDS.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 27

Preferred polypeptide encoded by this gene comprise the following amino acid sequence: NTNQREALQYAKNFQPFALNHQKDIQVLMGSLVYLRQGIENSPYVHL LDANQWADICDIFTRDACALLGLSVESPLSVSFSAGCVALPALINIKAVIEQRQC TGVWNQKDELPIEVDLGKKCWYHSIFACPILRQQTTDNNPPMKLVCGHIISRD ALNKMFNGSKLKCPYCPMEQSPGDAKQIFF (SEQ ID NO:276). Polynucleotides encoding such polypeptides are also provided as are complementary polynucleotides thereto.

This gene is expressed primarily in liver (adult and fetal) and spleen tissue, and to a lesser extent in placenta, T helper cells, kidney tumor, ovarian tumor, melanocytes and fetal heart.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune and developmental diseases and disorders and liver diseases such as liver cancer. Similarly, polypeptides and antibodies directed to these

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polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, circulatory and hematopoietic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., liver, spleen, placenta, blood cells, kidney, ovary and other reproductive tissue, melanocytes, and heart, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that the protein products of this gene are useful for study, diagnosis and treatment of growth, hematopoietic and immune system disorders particularly related to the liver.

15 FEATURES OF PROTEIN ENCODED BY GENE NO: 28

The translation product of this gene shares sequence homology with prostaglandin transporter which is thought to be important in metabolic and endocrine disorders. See, for example, Gastroenterology Oct:109(4):1274-1282 (1995). Preferred polypeptides encoded by this gene comprise the following amino acid sequence: SYLSACFAGCNSTNLTGCACLTTVPAENATVVPGKCPSPGCQEAFLTFLCVMCI CSLIGAMARHP (SEQ ID NO:277); and/or PSVIILIRTVSPELKSYALGVLFLLLRL LGFIPPPLIFGAGIDSTCLFWSTFCGEQGACVLYDNVVYRYLYVSIAIALKSFAFI (SEQ ID NO:278).

This gene is expressed primarily in hematopoietic and brain tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, metabolic, immune and endocrine diseases and disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the metabolic, immune and endocrine systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., endocrine tissue, hematopoietic tissue, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to

the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to prostaglandin (and anion) transporter indicates that polynucleotides and polypeptides corresponding to this gene are useful for study. diagnosis and treatment of endocrine, metabolic, immune and kidney disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 29

This gene is expressed primarily in early stage human lung.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, growth and respiratory disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the developmental and respiratory systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., pulmonary tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:153 as residues: Val-50 to Trp-55.

The tissue distribution indicates that the protein products of this gene are useful for study, diagnosis and treatment of respiratory and growth diseases and disorders.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 30

The translation product of this gene shares sequence homology with human DNA helicase which is thought to be important in accurate and complete DNA replication in creation of new cells. Preferred polypeptides encoded by this gene comprise the following amino acid sequence: QSLFTRFVRVGVPTVDLDAQGRARA SLCXXYNWRYKNLGNLPHVC_LPEFSTANAGLLYDFQLINVEDFQGVGESEPN PYFYQNLGEAEYVVALFMYMCLLGYPADKISILTTYNGQKHLIRDIINRRCGNN PLIGRPNKVTTVDRFQGQQNDYILLSLVRTRAVGHLRDVRRLVVAMSRAR (SEQ ID NO:279); and/or LVKEAKIIAMTCTHAALKRHDLVKLGFKYDNILMEE AAQILEIETFIPLLLQNPQDGFSRLKRWIMIGDHHQLPPVI (SEQ ID NO:280).

This gene is expressed primarily in testes tumor and to a lesser extent in adrenal

gland tumor and placenta.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cancers and endocrine/growth disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the endocrine, developmental, and reproductive systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., testes and other reproductive tissue, adrenal gland, and placenta, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to DNA helicase indicates that the protein products of this gene are useful for study, treatment, and diagnosis of many cancer types, including testicular cancer; as well as disorders involving endocrine function and normal growth and development.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 31

The translation product of this gene shares sequence homology with BID-apoptotic death gene (mouse). Genbank accession no. PID g1669514, which is thought to be important in programmed cell death.

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This gene is expressed primarily in jurkat membrane bound polysomes and activated neutrophils and to a lesser extent in endothelial cells and human cerebellum.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cancers and other proliferative disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., blood cells, endothelium, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma,

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urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:155 as residues: Glu-4 to Leu-11, Cys-28 to Arg-35, Gln-50 to His-66, Glu-73 to Gln-79, Gly-94 to Ser-100, Arg-114 to Asp-126, Pro-139 to Lys-146.

The tissue distribution and homology to BID-apoptotic death gene indicates that the protein products of this gene are useful for study of cell death, and treatment and diagnosis of proliferative disorders and cancers. Apoptosis - programmed cell death - is a physiological mechanism involved in the deletion of peripheral T lymphocytes of the immune system, and its dysregulation can lead to a number of different pathogenic processes. Diseases associated with increased cell survival, or the inhibition of apoptosis, include cancers (such as follicular lymphomas, carcinomas with p53 mutations, and hormone-dependent tumors, such as breast cancer, prostrate cancer, Kaposiís sarcoma and ovarian cancer); autoimmune disorders (such as systemic lupus erythematosus and immune-related glomerulonephritis rheumatoid arthritis) and viral infections (such as herpes viruses, pox viruses and adenoviruses), inflammation; graft vs. host disease, acute graft rejection, and chronic graft rejection. Diseases associated with increased apoptosis include AIDS; neurodegenerative disorders (such as Alzheimer's disease, Parkinson's disease, Amyotrophic lateral sclerosis, Retinitis pigmentosa, Cerebellar degeneration); myelodysplastic syndromes (such as aplastic anemia), ischemic injury (such as that caused by myocardial infarction, stroke and reperfusion injury), toxin-induced liver disease (such as that caused by alcohol), septic shock, cachexia and anorexia. Thus, the invention provides a method of enhancing apoptosis in an individual by treating the individual with a polypeptide encoded by this gene.

FEATURES OF PROTEIN ENCODED BY GENE NO: 32

The translation product of this gene shares sequence homology with human fructose transporter which is thought to be important in normal metabolic function and activity.

This gene is expressed primarily in T-cell lymphoma.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are

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not limited to, leukemia and other cancers, and metabolic disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the hematopoietic, lymph and metabolic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:156 as residues: Pro-22 to Gly-48, Ser-54 to Pro-61.

The tissue distribution indicates that the protein products of this gene are useful for study of mechanisms leading to cancer, treatment and diagnosis of cancerous and pre-cancerous conditions; as well as the study and treatment of various metabolic diseases and disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 33

This gene is expressed primarily in human meningima.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, inflammation and other disorders of the CNS. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the CNS and immune systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., meningima and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:157 as residues: Asn-23 to Pro-31.

The tissue distribution indicates that the protein products of this gene are useful for study, diagnosis and treatment of disorders of the CNS and inflammatory responses.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 34

This gene is expressed primarily in activated monocytes and wound healing tissues and to a lesser extent in fetal epithelium.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune and inflammatory disorders and wound healing and tissue repair dysfunctions. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, epithelial and gastrointestinal systems, and healing wounds, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., monocytes and other blood cells, and epithelium, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:158 as residues: Ala-28 to Ala-33, Gly-35 to Glu-45.

The tissue distribution indicates that the protein products of this gene are useful for diagnosis, study and treatment of immune and inflammatory disorders and wound healing dysfunctions.

25 FEATURES OF PROTEIN ENCODED BY GENE NO: 35

This gene is expressed primarily in human osteosarcoma and prostate cancer.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, skeletal and neoplastic conditions such as bone and prostate cancer. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and skeletal systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., bone, and prostate, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial

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fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:159 as residues: Ser-14 to Gly-22, Leu-37 to Gln-43.

The tissue distribution indicates that the protein products of this gene are useful for diagnosis and treatment of skeletal disorders and cancer.

FEATURES OF PROTEIN ENCODED BY GENE NO: 36

This gene encodes a protein which is highly homologous to a protein called congenital heart disease protein 5, presumably implicated in congenital heart disease (see Genbank PID g2810996).

This gene is expressed primarily in Hodgkin's lymphoma, erythroleukemia cells, and TNF activated synovial fibroblasts, to a lesser extent in ovarian cancer, cerebellum, spleen, fetal liver and placenta and finally to a lesser extent in various other mesenchymal tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cancer, immune, hematopoietic and cardiovascular disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, hematopoietic and cardiovascular systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., heart and other cardiovascular tissue, lymphoid tissue, blood cells, bone marrow, ovary and other reproductive tissue, brain and other tissue of the nervous system, spleen, liver, and mesenchymal tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:160 as residues: Lys-41 to Met-49, Gln-54 to Glu-59, Glu-76 to Thr-88.

The homology of this gene and translation product to congenital heart disease protein 5 indicates a role for this protein in the diagnosis, prognosis and/or treatment of

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heart disease or other cardiovascular related disorders. In addition, predominant expression in cells associated with the immune and hematopoetic system indicates a role for this protein in the treatment, diagnosis and/or prognosis of immune and autoimmune diseases, such as lupus, transplant rejection, allergic reactions, arthritis, asthma, immunodeficiency diseases, leukemia, AIDS, thymus disorders such as Graves Disease, lymphocytic thyroiditis, hyperthyroidism and hypothyroidism, graft versus host reaction, graft versus host disease, transplant rejection, myelogenous leukemia, bone marrow fibrosis, and myeloproliferative disease. The protein could also be used to enhance or protect proliferation, differentiation and functional activation of hematopoietic progenitor cells such as bone marrow cells, which could be useful for cancer patients undergoing chemotherapy or patients undergoing bone marrow transplantation. The protein may also be useful to increase the proliferation of peripheral blood leukocytes, which could be useful in the combat of a range of hematopoietic disorders including immunodeficiency diseases, leukemia, and septicemia.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 37

This gene is expressed primarily in ovarian cancer.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, urogenital neoplasias. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the reproductive system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., ovary and other reproductive tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:161 as residues: Asn-22 to Asn-27.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for study, diagnosis and treatment of ovarian and other tumors.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 38

The translation product of this gene shares sequence homology with zinc finger proteins.

This gene is expressed primarily in various fetal, cancer, and endothelial lines.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune and growth disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the cardiovascular system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., fetal tissue, and endothelial cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that the protein products of this gene are useful for study, diagnosis and treatment of immune and developmental conditions and cancer.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 39

This gene is expressed primarily in fetal, infant, and adult brain and to a lesser extent in other brain and endocrine organs and blastomas.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, brain tumors and neurodegenerative conditions. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the nervous and endocrine systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., brain and other tissue of the nervous system, endocrine tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the

disorder.

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The tissue distribution indicates that the protein products of this gene are useful for the study, diagnosis and treatment of brain cancer and other neurological disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 40

The translation product of this gene shares sequence homology with vesicular glycoproteins and lectins. Preferred polypeptides encoded by this gene comprise the following amino acid sequence: DTYPNEEKQQERVFPXXSAMVNNGSLSYDHER DGRPTELGGCXAIVRNLHYDTFLVIRYVKRHLTIMMDIDGKHEWRDCIEVPGV RLPRGYYFGTSSITGDLSDNHDVISLKLFELTVERTPEEE (SEQ ID NO:281); and/or LKREHSLSKPYQGVGTGSSSLWNLMGNAMVMTQYIRLTPDMQSKQGA LWNRVPCFLRDWELQVHFKIHGQGKKNLHGDGLAIWYT (SEQ ID NO:282).

This gene is expressed primarily in infant brain and to a lesser extent in various normal and transformed neural, endocrine, and immune organs.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, neurological and neurodevelopmental conditions. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the nervous and hormonal systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., brain and other tissue of the nervous system, endocrine tissue, and tissue and cells of the immune system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:164 as residues: Pro-64 to Gly-71, Gly-94 to Leu-100, Thr-110 to Pro-116, Thr-135 to Arg-145, Glu-164 to Glu-171, Asp-204 to Asp-211, Arg-253 to His-261, Asn-312 to Tyr-323.

The tissue distribution indicates that the protein products of this gene are useful for the study, diagnosis and treatment of mental retardation and other neurological disorders and neoplasias.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 41

This gene displays homology to the glycosyltransferase family, which catalyze the addition of sialic acids to carbohydrate groups which are present on glycoproteins.

This gene is expressed primarily in smooth muscle and to a lesser extent in pineal gland, fetal liver, and infant brain.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, gastrointestinal injury, inflammatory and neurodegenerative conditions. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and nervous systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., smooth muscle, pineal gland, liver, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:165 as residues: Ser-12 to Trp-21, Arg-24 to Pro-32, Asp-73 to Lys-82, Lys-90 to Ala-97.

The tissue distribution indicates that the protein products of this gene are useful for the study, diagnosis and treatment of neurodegenerative and growth disorders and gastrointestinal repair.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 42

The translation product of this gene shares sequence similarity with metallothionein polypeptides. See, for example, Proc. Natl. Acad. Sci. U S A 1992 Jul 15:89(14):6333-6337. Metallothioneins are believed to inhibit neuronal survival among other biological functions. Based on the sequence similarity (especially the conserved cysteine motifs characteristic of the metallothionein family) the translation product of this gene is expected to share certain biological activities with other members of the metallothionein polypeptide family. Preferred polypeptides encoded by this gene comprise the following amino acid sequence: PGTLQCSALHHDPGCANCSRFCRD CSPPACQC (SEQ ID NO:283).

This gene is expressed exclusively in placenta and fetal liver.

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Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, hematopoietic and immune disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the reproductive and immune systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., placenta, liver, brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to metallothionien indicates that the protein products of this gene are useful for diagnosis and treatment of immune and hematopoietic system disorders and neurological diseases, especially in fetal development.

20 FEATURES OF PROTEIN ENCODED BY GENE NO: 43

Preferred polypeptides encoded by this gene comprise the following amino acid sequence: FLYDVLMXHEAVMRTHQIQLPDPEFPS (SEQ ID NO:284).

This gene is expressed primarily in T-cells and synovial tissue.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune system disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., synovial tissue, and T-cells and other blood cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

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The tissue distribution indicates that the protein products of this gene are useful for treatment and diagnosis of disorders of the immune system.

FEATURES OF PROTEIN ENCODED BY GENE NO: 44

The translation product of this gene shares sequence similarity with several methyltransferases (e.g., see Genbank gil1065505).

This gene is expressed primarily in ovary, thymus, infant adrenal gland, tissues of the nervous system and the hematopoietic tissue, and to a lesser extent in adipose tissue and many other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, disorders of the reproductive system, the endocrine system, the hematopoietic system and the CNS. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, endocrine, CNS and reproductive system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., ovary and other reproductive tissue, thymus, adrenal gland, brain and other tissue of the nervous system, hematopoietic tissue, and adipose tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:168 as residues: Ser-3 to Gly-12, Asp-19 to Arg-31, Tyr-70 to Tyr-77, Asn-130 to Lys-140, Pro-165 to Gln-170, Pro-192 to Lys-199, Leu-216 to Glu-227, Glu-254 to Phe-281.

The tissue distribution and homology to methyltransferase indicates that the protein products of this gene are useful for diagnosis and treatment of disorders of the CNS, the hematopoietic system and reproductive organs and tissues. For example, the abundant expression in the ovary may indicate that the gene product can be used as a hormone with either systemic or reproductive functions; as growth factors for germ cell maintenance and in vitro culture; as a fertility control agent; remedy for sexual dysfunction or sex development disorders; diagnostics/treatment for ovarian tumors, such as serous adenocarcinoma, dysgerminoma, embryonal carcinoma,

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choriocarcinoma, teratoma, etc: The expression in thymus may indicate its utilities in T-cell development and thus its applications in immune related medical conditions, such as infection, allergy, immune deficiency, tissue/organ transplantation, etc.

5 FEATURES OF PROTEIN ENCODED BY GENE NO: 45

The translation product of this gene shares sequence homology with cytochrome C oxidase which is thought to be important in metabolic function of cells. This gene has now recently been published as estrogen response gene. See Genbank accession no. AB007618 and Mol. Cell. Biol. 18 (1), 442-449 (1998). See also J Immunol. Mar 1:154(5): 2384-2392 (1995), where the mouse homologue was published and implicated in siliocis.

This gene is expressed primarily in adipose tissue, kidney and fetal brain and to a lesser extent in several other tissues and organs.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, metabolic diseases involving especially adipose tissue, brain and kidney. Similarly, polypeptides and antibodies directed to these polypeptides are useful inproviding immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the CNS and vascular system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., adipose tissue, kidney, brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:169 as residues: Thr-5 to Ser-14.

The tissue distribution and homology to cytochrome C oxidase, estrogen response gene product and siliocis related gene product indicates that the protein products of this gene are useful for diagnosis and treatment of metabolic disorders in the CNS, adipose tissue and kidney, particularly siliocis.

FEATURES OF PROTEIN ENCODED BY GENE NO: 46

The translation product of this gene shares sequence homology with reticulocalbin. See, for example, J. Biochem. 117 (5), 1113-1119 (1995). Based on the

sequence similarity, the translation product of this gene is expected to share certain biological activities with reticulocalbin, e.g., Ca++ binding activities. This gene product is sometimes hereinafter referred to as "Reticulocalbin-2".

This gene is expressed primarily in breast, endothelial cells, synovial, heart and smooth muscle cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, diseases of the breast, vascular and skeletal/cardiac muscular system. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the breast, vascular and skeleto-muscular system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., mammary tissue, endothelial cells, synovial tissue, heart and other cardiovascular tissue, and smooth muscle, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:170 as residues: Gly-16 to Arg-32, Ala-42 to Asn-50, Glu-66 to Gln-76, Arg-85 to Gly-94, Thr-108 to Asp-115, Trp-121 to Gly-130. Leu-137 to His-144, Glu-155 to Lys-161, Asp-175 to Ser-180, Glu-209 to Gly-217, Glu-232 to Glu-237, Thr-243 to Asp-261, Glu-287 to Arg-295.

The tissue distribution indicates that the protein products of this gene are useful for diagnosis and treatment of diseases of the vascular and skeletal/cardiac muscular system. The homology of the gene with reticulocalbin indicates its biological function in regulating calcium store, a particularly important function in muscular cell types. The gene expression in the heart may indicate its utilities in diagnosis and remedy in heart failure, ischemic heart diseases, cardiomyopathy, hypertension, arrhythmia, etc. The abundant expression in the breast may indicate its applications in breast neoplasia and breast cancers, such as fibroadenoma, papillary carcinoma, ductal carcinoma, Pagetís disease, medullary carcinoma, mucinous carcinoma, tubular carcinoma, secretory carcinoma and apocrine carcinoma; juvenile hypertrophy and gynecomastia, mastitis and abscess, duct ectasia, fat necrosis and fibrocystic diseases, etc.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 47

The translation product of this gene shares weak sequence homology with H+-transporting ATP synthase which is thought to be important in cell metabolism or signal transduction.

This gene is expressed only in testis.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of some types of diseases and conditions. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the brain and hematopoietic tissues, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., testes and other reproductive tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

Since only one out of about a million expressed sequence tag is found in testes indicates that its expression is low and selectively in testes. Since some of the genes only expressed in testes are usually expressed in brain or in certain induced hematopoietic cells/tissues, it is speculated that this gene to be expressed in brain or hematopoietic cells/tissues and is useful for diagnosis and treatment of disorders these systems.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 48

The translation product of this gene shares sequence homology with human polymeric immunoglobulin receptor (accession No.X73079) which is thought to be important in antibody recognition and immune defenses. In one embodiment, polypeptides of the invention comprise the sequence GWYWCG (SEQ ID NO:285). Polynucleotides encoding these polypeptides are also encompassed by the invention.

This gene is expressed primarily in placenta and to a lesser extent in corpus callosum and fetal liver and spleen.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are

not limited to, disorders of the immune system, e.g. autoimmune diseases and immunodeficiency. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., placenta, liver, and spleen, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:172 as residues: Tyr-37 to Cys-49, Gly-51 to Tyr-56, Lys-88 to Trp-93, Leu-130 to Glu-136.

The tissue distribution and homology to human polymeric immunoglobulin receptor indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of immune disorders, e.g. autoimmune diseases and immunodeficiencies.

FEATURES OF PROTEIN ENCODED BY GENE NO: 49

This gene is expressed in thymus.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune disorder. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., thymus and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of immune disorders, e.g. autoimmunity and immunodeficiency.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 50

Preferred polypeptide encoded by this gene comprise the following amino acid sequence: MKVGARIRVKMSVNKAHPVVSTHWRWPAEWPQMFLHLAQEPRTE VKSRPLGLAGFIRQDSKTRKPLEQETIMSAADTALWPYGHGNREHQENELQKY LQYKDMHLLDSGQSLGHTHTLQGSHNLTALNI (SEQ ID NO:286). Polynucleotides encoding this polypeptide are also provided as are complementary polynucleotides thereto.

This gene is expressed primarily in adrenal gland, pituitary, T helper cells, and breast cells and to a lesser extent in a wide variety of tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of the some diseases and conditions. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and endocrine systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., adrenal gland, pituitary, T-cells and other blood cells, and mammary tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:174 as residues: Gln-39 to Ser-47, Arg-57 to Glu-67, Tyr-82 to Gln-95.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of a wide range of disorders, such as immune and endocrine disorders.

30 FEATURES OF PROTEIN ENCODED BY GENE NO: 51

The translation product of this gene shares sequence homology with human Sop2p-like protein which is important in cytoskeleton structure. In one embodiment, polypeptides of the invention comprise the sequence SLHKNSVSQISVLSGGKAKCS QFCTTGMDGGMSIWDVKSLESALKDLKI (SEQ ID NO:287). Polynucleotides encoding this polypeptide are also encompassed by the invention. This gene maps to chromosome 7. Therefore, polynucleotides of the invention can be used in linkage

analysis as a marker for chromosome 7.

This gene is expressed primarily in immune and hematopoietic tissues/cells and to a lesser extent in other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immunological and hematopoietic disorders and inflammation. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and hematopoietic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., immune and hematopoietic tissue/cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:175 as residues: Lys-49 to Gln-54, Ala-61 to Arg-66, Lys-82 to Lys-87, Glu-126 to Val-133, His-136 to Ile-141, Glu-175 to Ser-187, Asp-286 to Leu-296, Ala-298 to Ser-310.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of immunological, hematopoietic, and inflammatory disorders, e.g., immunodeficiency, autoimmunity, inflammation.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 52

The translation product of this gene shares sequence homology with *Caenorhabditis elegans* R53.5 gene encoding a putative secreted protein without known function.

This gene is expressed primarily in endothelial cells, brain and several highly vascularized, and tumor tissues and to a lesser extent in other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, aberrant angiogensis and tumorigenesis. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes

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for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the vascular and brain system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., endothelial cells, brain and other tissue of the nervous system, and vascular tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:176 as residues: Thr-43 to Asn-60, Thr-106 to Phe-115, Asp-122 to Arg-133, Arg-186 to Asp-192, Leu-211 to Lys-216.

The tissue distribution and homology to a *C. elegans* secreted protein indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis or treatment of disorders in vascular or brain system, e.g. aberrant angiogenesis, ischemia, neurodegeneration, etc.

FEATURES OF PROTEIN ENCODED BY GENE NO: 53

In one embodiment, polypeptides of the invention comprise the sequence EASKSSHAGLDLFSVAACHRF (SEQ ID NO:288). Polynucleotides encoding this polypeptide are also encompassed by the invention.

This gene is expressed primarily in T-cells and to a lesser extent in brain.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, lymphocytic disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the lymphoid system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:177 as residues: Pro-3 to Thr-8, Arg-37 to Asp-46.

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The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis, treatment, and cure of lymphocytic disorders.

5 FEATURES OF PROTEIN ENCODED BY GENE NO: 54

The translation product of this gene shares sequence homology with secreted cartilage matrix protein, a major component of the extracellular matrix of nonarticular cartilage which is thought to be important in cartilage structure. In specific embodiments, polypeptides of the invention comprise the sequence: RCKKCTEGPI DLVFVIDGSKSLGEENFEVVKQF (SEQ ID NO:297); VTGIIDSLTISPKAARVGL LQYSTQVH (SEQ ID NO:290); TEFTLRNFNSAKDMKKAVAHMKYM (SEQ ID NO:291); GKGSMTGLALKHMFERSFTQGEGARPF (SEQ ID NO:292); STRVP RAAIVFTDGRAQDDVSEWASKAKANGITMYAVGVGKAIE (SEQ ID NO:293); EELQEIASEPTNKHLFYAEDFSTMDEISEKLKKGICEALEDS (SEQ ID NO:294); TQRLEEMTQRM (SEQ ID NO:295); PQGCPEQPLH (SEQ ID NO:296); and/or YMGKGSMTGLALKHMFERSFT (SEQ ID NO:289). Polynucleotides encoding these polypeptides are also encompassed by the invention.

This gene is expressed primarily in placenta, infant brain, prostate, fetal lung and to a lesser extent in endometrium and fetal tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, abnormal placenta and pregnancy, disorder and injury in brain, prostate, and vasculature. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the reproduction, neuronal, and vascular systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., placenta, brain and other tissue of the nervous system, prostate, lung and endometrium, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to cartilage matrix protein indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis,

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treatment, and cure of abnormalities in placenta and pregnancy, disorder and injury in brain, prostate, and vasculature.

FEATURES OF PROTEIN ENCODED BY GENE NO: 55

The translation product of this gene is the human ortholog of bovine and hamster CII-3, a succinate-ubiquinone oxidoreductase complex II membrane-intrinsic subunit, which is thought to be important in mitochondrial electron transport chain during metabolism. In specific embodiments, the polypeptides of the invention compriseMAALLLRHVGRHCLRAHFSPQLCIRNAVPLGTTAKEEMERFWNKNIG SNRPLSPHITIYS (SEQ ID NO:298); VFPLMYHTWNGIRHLMWDLGKGLKIPQL YQSG (SEQ ID NO:299); MAALLLRHVGRHCLRAH (SEQ ID NO:300); VKSLCL GPALIHTAKFAL (SEQ ID NO:301); VFPLMYHTWNGIRHLMWDLGKGL (SEQ ID NO:302).

This gene is expressed in 8-week old early stage human.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, metabolism disorder. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the [insert system where a related disease state is likely, e.g., immune], expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis, treatment, and cure of metabolism disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 56

This gene is expressed primarily in umbilical vein endothelial cells, human ovarian tumor cells, human meningima cells, and human Jurkat membrane bound polysomes. In specific embodiments, polypeptides of the invention comprise the amino acid sequence: RVWDVRPFAPKERCVKIFQGNV (SEQ ID NO:303); HNFEKNLL

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RCSWSPDGSKIAAGSADRFVYV (SEQ ID NO:304); and/or WDTTSRRILYKLPG HAGSINEVAFHPDEPI (SEQ ID NO:305). Polynucleotides encoding these polypeptides are also encompassed by the invention.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, inflammation, immune and cardiovascular disorders and urogenital neoplasias. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of these tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, neurological, urogenital, reproductive system and vascular systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., blood cells, cells, endothelial cells, ovary and other reproductive tissue, meningima, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:143 as residues: Phe-71 to Arg-76, Pro-82 to His-87, Glu-103 to Ala-111.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the diagnosis and treatment of immune disorders including: leukemias, lymphomas, auto-immune, immuno-supressive (e.g. transplantation) and immunodeficiencies (e.g. AIDS) and hematopoietic disorders. In addition, expression in ovarian tumor cells suggests that polynucleotides and polypeptides corresponding to this gene are useful for study, diagnosis, and treatment of ovarian tumors, and other tumors and neoplasias. Further, endothelial cell expression suggests a role in cadiovascular or respiratory/pulmonary disorders or infections (athsma, pulmonary edema, pneumonia).

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FEATURES OF PROTEIN ENCODED BY GENE NO: 57

The translation product of this gene shares sequence homology with type I collagen. In specific embodiments, the polypeptides of the invention comprise the sequence: GRIPAPAPSVPAGPDSR (SEQ ID NO:309); VRGRTVLRPGLDAEPE LSPE (SEQ ID NO:306); EQRVLERKLKKERKKEERQ (SEQ ID NO:307); ARRSG

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AELAWDYLCRWAQKHKNWRFQKTRQTWLLLHMYDSDKVPDEHFSTLLAYLE GLQGR (SEQ ID NO:255); and/or RLREAGLVAQHPP (SEQ ID NO:308).

Polynucleotides encoding these polypeptides are also encompassed by the invention.

This gene is expressed primarily in epididymus, prostate cell line (LNCAP), and pituitary gland; and to a lesser extent in many other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, abnormalities of the epididymus, prostate (especially prostate cancer), and pituitary gland. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the male reproductive system and neuroendocrine system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., epididymus and other reproductive tissue, prostate, and pituitary gland, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution and homology to type I collagen, indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of abnormalities of the epididymus, prostate (especially prostate cancer), and pituitary gland.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 58

This gene is expressed primarily in the frontal cortex of the brain from a schizophrenic individual.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, schizophrenia. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the nervous system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., brain and other tissue of

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the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum. plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of schizophrenia.

FEATURES OF PROTEIN ENCODED BY GENE NO: 59

The polypeptide encoded by Gene 59 is homologous to human surface 4 integral membrane protein. In specific embodiments, the polypeptides of the invention comprise the sequence: TGCVLVLSRNFVQYACFGLFGIIALQTIAYSILWDLKF LMRN (SEQ ID NO:310); SRSEGKSMFAGVPTMRESSPKQYMQLGGRVLLV LMFMTLLHFDASFFSIVQNIVG (SEQ IDNO:311); GTAEDFADQFLRVTKQYLP HVARLCLISTFLEDGIRMFQWSEQRDYIDTTWNCGYLLAS (SEQ ID NO:312); LMRNESRS (SEQ ID NO:314); ASFLLSRTSWGTA (SEQ ID NO:315); and/or ASFLLSRTSWGTALMIL (SEQ ID NO:313). Polynucleotides encoding these polypeptides are also encompassed by the invention.

This gene is expressed primarily in Hodgkin's lymphoma and lung; and to a lesser extent in many other human tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, Hodgkin's lymphorna, tumors or other abnormalities of the lung. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and respiratory systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., lymphoid tissue, and pulmonary tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:183 as residues: Met-20 to Trp-27.

The tissue distribution indicates that polynucleotides and polypeptides

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corresponding to this gene are useful for diagnosis and treatment of Hodgkin's lymphoma, tumors or other abnormalities of the lung.

FEATURES OF PROTEIN ENCODED BY GENE NO: 60

This gene is expressed primarily in bone cancer and stomach cancer, and to a lesser extent in many other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, bone cancer and stomach cancer. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the bone, and the stomach, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., bone, and stomach, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of bone cancer and stomach cancer and possibly other cancers.

FEATURES OF PROTEIN ENCODED BY GENE NO: 61

This gene is expressed primarily in epididymus, and lymph node of breast cancer, and to a lesser extent in many other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, abnormalities of the epididymus, and breast cancer. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the epididymus and breast, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., epididymus and other reproductive tissue, lymphoid tissue, and mammary tissue, and cancerous and wounded tissues) or bodily fluids

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(e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:185 as residues: Arg-57 to Ser-65.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of abnormalities of the epididymus, and breast cancer.

10 FEATURES OF PROTEIN ENCODED BY GENE NO: 62

The translation product of this gene appears to be the human homolog of bovine NADH dehydrogenase which is thought to be important in cellular metabolism. In specific embodiments, the polypeptides of the invention comprise the amino acid sequence: SMSALTRLASFARVGGRLFRSGCARTAGDGGVRHAGGGVHIEPRY RQFPQLTRSQVFQSEFFSGLMWFWILWRFWHDSEEVLGHFPYPDPSQWTDEEL GIPPDDED (SEQ ID NO:323), or fragments thereof. Polynucleotides encoding this polypeptide are also encompassed by the invention.

This gene is expressed in larynx tumor, lymph node, brain amygdala, human cardiomyopathy, and retina.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, diseases affecting cellular metabolism. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the nervous system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., larynx, lymphoid tissue, brain and other tissue of the nervous system, heart and cardiovascular tissue, and retina, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:208 as residues: Pro-27 to Gln-32, Arg-42 to Glu-51.

The tissue distribution and homology to NADH dehydrogenase indicates that

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polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of diseases involving cellular metabolism.

FEATURES OF PROTEIN ENCODED BY GENE NO: 63

This gene is expressed primarily in amygdala, and to a lesser extent in many other tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, abnormalities of the amygdala. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the amygdala, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., amygdala, and lymphoid tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:187 as residues: Gln-17 to Glu-29, Pro-41 to Phe-46, Ser-59 to Ile-70, Thr-97 to Leu-105.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for diagnosis and treatment of abnormalities of amygdala.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 64

This gene is expressed primarily in female bladder, and to a lesser extent in chronic synovitis and hemangiopericytoma.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, bladder cancer. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the urinary tract, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., bladder, synovial tissue, and

vascular tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:188 as residues: Pro-2 to Gln-7, Pro-27 to Phe-34.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatments of defects of the urinary tract, especially bladder cancer.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 65

This gene is expressed primarily in fetal spleen, and to a lesser extent in hemangiopericytoma, thymus, and synovial sarcoma.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, defects of immune of hematopoietic systems. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune of hematopoietic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., spleen, vascular tissue, thymus, blood cells, and synovial tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The protein product of this gene is useful for treatment of defects of the immune or hematopoietic systems, because of the gene's expression in thymus and spleen.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 66

This gene is expressed primarily in human pituitary and to a lesser extent in placenta and fetal lung.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are

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not limited to, endocrine growth disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the endocrine system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., pituitary and other endocrine tissue, placenta, and pulmonary tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:190 as residues: Val-38 to Asn-44, Gly-53 to Ser-65.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment of growth disorders related to pituitary dysfunction.

FEATURES OF PROTEIN ENCODED BY GENE NO: 67

The translation product of this gene shares sequence homology with a Caenorhabditis elegans gene of unknown function. In specific embodiments, the polypeptides of the invention comprise the sequence: DPRRPNKVLRYKPPPSE CNPALDDPTP (SEQ ID NO:317); DYMNLLGMIFSMCGLMLKLKWCAWVA VYCS (SEQ ID NO:318); FISFANSRSSEDTKQMMSSF (SEQ ID NO:316); and/or MLSISAVVMSYLQNPQPMTPPW (SEQ ID NO:319). Polynucleotides encoding these polypeptides are also encompassed by the invention.

This gene is expressed primarily in primary breast cancer and lymph node breast cancer and to a lesser extent in adult brain, lung cancer, colon cancer, epithelioid-sarcoma, and Caco-2 cell line.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cancer. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the cancer and tumor tissues, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., mammary tissue, lymphoid tissue, brain and other tissue of the nervous system, lung, colon, and

epithelium, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:191 as residues: Asn-34 to Lys-42.

The tissue distribution in a variety of cancer tissues indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of a variety of cancer and tumor types.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 68

The translation product of this gene shares sequence homology with steroid membrane binding protein. The translation product of this gene has recently been published as progesterone binding protein. See Genbank AJ002030. Preferred polypeptides encoded by this gene comprise the following amino acid sequence: AAGDGDVKLGTLGSGSESSNDGGSESPGDAGAAAXGGGWAAAALALLTG GGE (SEQ ID NO:320).

This gene is expressed primarily in breast, and to a lesser extent in placenta and fetal tissue.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, breast cancer or developmental disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of breast or fetal tissues, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., mammary tissue, placenta, and fetal tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:192 as residues: Pro-43 to Asp-49, Gln-54 to Pro-64, Asp-110 to Asp-118, Lys-138 to Tyr-143, Pro-150 to Asp-170.

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The tissue distribution and homology to steroid membrane binding protein and to progesterone binding protein indicates that the protein products of this gene are

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useful for treatment of breast cancers, especially those caused by estrogen and progesterone binding.

FEATURES OF PROTEIN ENCODED BY GENE NO: 69

Preferred polypeptides encoded by this gene comprise the following amino acid sequence: AADNYGIPRACRNSARSYGAAWLLLXPAGSSRVEPTQDISISDQLGG QDVPVFRNLSLLVVGVGAVFSLLFHLGTRERRPHAXEPGEHTPLLAPATAQPL LLWKHWLREXAFYQVGILYMTTRLIVNLSQTYMAMYLTYSLHLPKKFIATIPLV MYLSGFLSSFLMKPINKCIGRN (SEQ ID NO:321).

This gene is expressed primarily in macrophage (GM-CSF treated), and to a lesser extent in monocytes and dendritic cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, inflammation and infection. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., macrophages and other blood cells, and dendritic cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that the protein products of this gene are useful for treatment of infection or inflammation or other events or defects involving the immune system.

FEATURES OF PROTEIN ENCODED BY GENE NO: 70

This gene is expressed primarily in adult brain and to a lesser extent in thyroid, 12 week old early stage human, and stromal cell TF274.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, neurological or neuro-endocrine diseases. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes

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for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the central nervous or endocrine systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., brain and other tissue of the nervous system, thyroid, and stromal cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:194 as residues: Pro-65 to Cys-71.

The tissue distribution indicates that the protein products of this gene are useful for treatment and diagnosis of neurological diseases or metabolic conditions involving the neuro-endocrine system.

FEATURES OF PROTEIN ENCODED BY GENE NO: 71

This gene is expressed in T-cell helper and to a lesser extent in adult brain and adult testes.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, immune disorders, meningitis or reproductive problems. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune, neural and reproductive systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, brain and other tissue of the nervous system, testes and other reproductive tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:195 as residues: Val-18 to Tyr-24, Ala-89 to Asp-99, Asp-104 to Ala-117, Leu-121 to Pro-136.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis immune and

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reproductive disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 72

The translated polypeptide of this contig has a high degree of identity with the Ob Receptor-Associated Protein deposited as GenBank Accession No. 2266638. No function has been determined for the Ob Receptor-Associated Protein, however it is expressed upon stimulation of the Ob Receptor by Leptin.

This gene is expressed in T-cells and to a lesser extent in endothelial and bone marrow cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a . biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, acute lymphoblastic leukemia, hematapoetic disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and hematapoetic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, endothelial cells, and bone marrow, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:196 as residues: Ser-61 to Trp-70.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of leukemia and other disorders of the primary immune system. In addition, since this gene appears to be related to the Ob Receptor-Related Protein, it is likely that this polypeptide is also involved in the Ob/Leptin signal transduction cascade. As a result, this protein may be of use in the molecular diagnosis and therapeutic intervention of obesity and related disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 73

The translation product of this contig has homology with furin, a protein thought to be a key endopeptidase in the constitutive secretory pathway. The identification and initial characterization of Furin was reported by Takahasi and

colleagues (Biochem Biophys Res Commun 1993 Sep 15;195(2):1019-1026).

This gene is expressed in neutrophils.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, diseases of the immune system such as allergies, wound healing and antigen recognition. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., neutrophils and other blood cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment of allergies or other immune disorders since neutrophils are an important part of an allergic response. Further, since this protein appears to be related to Furin, it can be used diagnostically and therapeutically to treat secretory protein processing disorders.

FEATURES OF PROTEIN ENCODED BY GENE NO: 74

This gene is expressed in the frontal cortex.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, of the motor activity and sensory functions that involve the central nervous system. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the central nervous system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene

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expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the detection and treatment of neural disorders that affect cognitive functions.

FEATURES OF PROTEIN ENCODED BY GENE NO: 75

The translation product of this gene shares sequence homology with inorganic pyrophophatase which is thought to be important in the catalysis the hydrolysis of diphosphate bonds, chiefly in nucleoside di- and triphosphates and essential enzymes that are important for controlling the cellular levels of inorganic pyrophosphate (PPi). The bovine homolog of this gene has been identified by Yang and Wensel (J. Biol. Chem. 267:24641-24647 (1992)).

This gene is expressed in osteoclastoma cells and to a lesser extent in epithelial cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, osteoporosis and other bone weakening diseases. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the skeletal system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., bone, and epithelial cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:199 as residues: Lys-22 to Tyr-28, Asp-64 to Lys-77, Pro-86 to Ile-91, Gln-99 to Pro-119, Tyr-169 to Asp-174, Lys-176 to Gly-181, Trp-189 to Asn-202, Lys-233 to Gly-239, Ser-250 to Asp-257.

The tissue distribution and homology to inorganic pyrophophatase indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of osteoporosis through the removal of bone by demineralization.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 76

The translation product of this gene shares exact sequence homology with ATP sulfurylase/APS kinase (GenBank Accession No. 2673862) which is thought to be important in biosynthesis of the activated sulfate donor, adenosine 3'-phosphate 5'-phosphosulfate, involves the sequential action of two enzyme activities: ATP sulfurylase, which catalyzes the formation of adenosine 5'-phosphosulfate (APS) from ATP and free sulfate, and APS kinase, which subsequently phosphorylates APS to produce adenosine 3'-phosphate 5'-phosphosulfate.

This gene is expressed in osteoclastoma cells and to a lesser extent in developmental tissues.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, antibiotic resistant bacterial infections, osteoarthritis and other auto immune diseases. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune or skeletal structure expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., bone, and developmental tissues, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:200 as residues: Asn-15 to Trp-20, Ser-36 to Gly-41, Pro-103 to Val-110, Pro-134 to Arg-143, Leu-173 to Arg-178, Ser-190 to Ala-197, His-314 to Arg-319, Arg-354 to Asn-362, Asp-391 to Arg-397, Glu-402 to Asp-409, Asp-434 to Leu-439, Glu-441 to Arg-446, Gly-455 to Asp-462, Pro-528 to His-541, Asn-566 to Arg-571, Tyr-574 to Glu-581, Thr-589 to Glu-603.

The tissue distribution and homology to ATP sulfurylase/APS kinase indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment or detection of autoimmune diseases.

FEATURES OF PROTEIN ENCODED BY GENE NO: 77

This polypeptide is identical to the SLP-76-associated protein reported by Musci and colleagues (J. Biol. Chem. 272 (18), 11674-11677 (1997)) and to the FYB protein

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reported by da Silva and coworkers (Proc. Natl. Acad. Sci. U.S.A. (1997) In press). These proteins have been reported to be novel T-cell Proteins which bind FYN and SLP-76 and regulate IL-2 production. Preferred polypeptides encoded by this genc comprise the following amino acid sequence: RITDNPEGKWLGRTARGSYGYIK TTAVEIXYDSLKLKKDSLGAPSRPIEDDQEVYDDVAEQDDISSHSQSGSGGIFPP PPDDDIYDGIEEEDADDGFPAPPKQLDMGDEVYDDVDTSDFPVSSAEMSQGTNV GKAKTEEKDLKKLKKQXKEXKDFRKKFKYDGEIRVLYSTKVTTSITSKKWGT RDLQVKPGESLEVIQTTDDTKVLCRNEEGKYGYVLRSYLADNDGEIYDDIADGC IYDND (SEQ ID NO:322).

This gene is expressed in CD34 positive cells (hematopoietic progenitor cells) and to a lesser extent in adult spleen derived from a chronic lymphocytic leukemia patient.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are 15 not limited to, chronic lymphocytic leukemia; hematopoietic disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune and 20 hematopoietic systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., T-cells and other blood cells, bone marrow, hematopoietic cells, and spleen, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily 25 fluid from an individual not having the disorder. Further, nucleic acids and polypeptides of the present invention are useful both diagnostically and therapeutically in the intervention of immune and other disorders in which the ability to alter IL-2 expression is desired. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:201 as residues: Ala-17 to Lys-37, Val-39 to Ser-45, Lys-59 to His-70, 30 Arg-90 to Leu-95, Lys-97 to Lys-107, Ser-117 to Leu-124, Phe-133 to Ser-138, Trp-146 to Leu-167, Pro-175 to Asn-185, Lys-190 to Ser-211, Pro-213 to Ser-222, His-230 to Pro-235, Pro-240 to Pro-246, Pro-253 to Gly-261, Leu-271 to Leu-303, Leu-305 to Leu-326, Lys-343 to Leu-349, Thr-363 to Leu-371, Arg-373 to Tyr-381, Tyr-35 391 to Leu-401, Pro-404 to Val-414, Ser-426 to Ser-432, Ile-448 to Ser-457, Gln-462 to Trp-468, Lys-477 to Ser-501, Asp-518 to Ser-523, Ala-541 to Gln-554.

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The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment of a variety of hematopoietic disorders. The noted expression of this gene in the hematopoietic progenitor cell compartment - as determined by its expression on CD34 positive hematopoietic stem and progenitor cells - indicates that it plays a critical role in the expansion or proliferation of hematopoietic stem/progenitor cells, as well as in the differentiation of the various blood cell lineages. Thus it could be useful in the reconstitution of the hematopoietic system of patients with leukemias and other hematopoietic diseases.

FEATURES OF PROTEIN ENCODED BY GENE NO: 78

This gene is homologous to heparin cofactor II (HCII) which is a 66-kDa plasma glycoprotein that inhibits thrombin rapidly in the presence of dermatan sulfate or heparin.

This gene is expressed in apoptotic and anergic T-cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, thrombopienia T-cell lymphomas; Hodgkin's lymphoma. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system - most notably the T-cell compartment, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, and lymphoid tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The homology to heparin cofactor II (HCII) and the tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment and diagnosis of hematopoietic disorders particularly in thrombopoesis, most notably of the T-cell compartment. This could include immune modulation, inflammation, immune surveillance, graft rejection, and autoimmunity.

35 FEATURES OF PROTEIN ENCODED BY GENE NO: 79

The translation product of this gene shares sequence homology with a mouse

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protein believed to represent an integral membrane protein.

This gene is expressed in fetal cochlea and epididymus and to a lesser extent in adult spleen and osteoclastoma.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, osteoclastoma; disorders of the inner ear; male fertility disorders. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the inner ear; male reproductive tract; bone; and immune systems, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., cochlea, epididymus and other reproductive tissue, spleen, and bone, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:203 as residues: Lys-13 to Gly-23, Cys-38 to Asp-43, Gly-48 to Trp-53, Cys-223 to Ile-237, Ile-240 to Ser-246.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for the treatment of hearing and fertility disorders. Likewise, it may have a role in the modulation of immune function and in the treatment of osteoporosis.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 80

The translation product of this gene shares sequence homology with reticulocalbin which is thought to be important in the binding of calcium, particularly within the endoplasmic reticulum.

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This gene is expressed in endothelial cells and stromal cells and to a lesser extent in osteoblasts, osteoclasts, and T-cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, osteoperosis; osteoclastomas; T-cell lymphomas; Hodgkin's disease. Similarly, polypeptides and antibodies directed to these polypeptides are useful in

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providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the vasculature, bone, and immune systems - particularly the T-cell compartments, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., endothelial cells, stromal cells, bone, T-cells and other blood cells, and lymphoid tissue, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:204 as residues: Lys-20 to Arg-27, Pro-32 to Asp-48, Leu-64 to Arg-72, Asp-108 to Lys-114, Glu-128 to Thr-133, Asp-139 to Phe-147, Thr-196 to Ala-204, Tyr-218 to Glu-228, Val-230 to Gln-236, Arg-241 to Lys-255, Glu-276 to Lys-287.

The tissue distribution and homology to reticulocalbin indicates that polynucleotides and polypeptides corresponding to this gene are useful for the diagnosis and treatment of bone disorders such as osteoporosis; the diagnosis and treatment of T-cell lymphomas and Hodgkin's lymphoma; and the treatment of diseases and defects of the vasculature, such as vascular leak syndrome and aberrant angiogenesis that accompanies tumor growth.

FEATURES OF PROTEIN ENCODED BY GENE NO: 81

The translation product of this gene shares sequence homology with a family of peptide transport genes - particularly the AtPTR2-B gene from *Arabidopsis* - which are thought to be important in the uptake of small peptides.

This gene is expressed in a number of fetal tissues, most notably lung, brain, cochlea, and liver/spleen, and to a lesser extent in osteoclastoma and endometrial tumors.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, osteoclastoma; endometrial tumors; cancer; leukemias. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the bone and endometrium, expression of this gene at significantly higher or lower levels may be

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routinely detected in certain tissues (e.g., fetal tissue, pulmonary tissue, bone, brain and other tissue of the nervous system, cochlea, liver, and spleen, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:205 as residues: Lys-186 to Asn-199, Pro-202 to Ala-207.

The tissue distribution and homology to peptide transport genes indicates that polynucleotides and polypeptides corresponding to this gene are useful for the control of cell proliferation, owing to its strong expression in fetal tissues undergoing active cell division, as well as its expression in a variety of tumors or cancers of adult tissues. Potentially, it may regulate the uptake of peptides that stimulate cell proliferation. This gene product may also be useful in stimulating the uptake of a variety of peptide-based drug compounds.

FEATURES OF PROTEIN ENCODED BY GENE NO: 82

This gene is expressed in fetal liver and spleen and to a lesser extent in endothelial cells.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, cancer and tumors of a hematopoietic and/or endothelial cell origin; leukemias. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system and/or vasculature, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., liver, spleen, endothelial cells, vascular tissue, and tissue and cells of the immune system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:206 as residues: Met-1 to Asp-9, Arg-66 to Gly-76, Asp-164 to Arg-171.

The tissue distribution indicates that polynucleotides and polypeptides

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corresponding to this gene are useful for the treatment of disorders of the immune system. Expression of this gene product in both fetal liver/spleen and endothelial cells indicates that it may be expressed in the hemangioblast, the progenitor cell for both the immune system and the vasculature. Thus, it is most likely expressed in hematopoietic stem cells, and may be useful for the expansion of hematopoietic stem and progenitor cells in conjunction with cancer treatment for a variety of leukemias.

FEATURES OF PROTEIN ENCODED BY GENE NO: 84

The translation product of this gene shares sequence homology with NADH dehydrogenase which is thought to be important in cellular metabolism.

This gene is expressed in fetal dura mater and to a lesser extent in T-cells and hypothalamus.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, diseases affecting cellular metabolism. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the nervous system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., fetal tissue, T-cells and other blood cells, and brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:208 as residues: Pro-27 to Gln-32, Arg-42 to Glu-51.

The tissue distribution and homology to NADH dehydrogenase indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of diseases involving cellular metabolism.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 85

The translation product of this gene shares sequence homology with I-TRAF, a novel TNF receptor associated factor (TRAF)-interacting protein that regulates TNF receptor-mediated signal transduction. This protein is thought to be important in regulating the cellular response to tumor necrosis factor (TNF), which is an important mediator of inflammation.

This gene is expressed in endothelial cells and to a lesser extent in glioblastoma and osteoblastoma.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, inflammation; glioblastoma and osteoblastoma. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., endothelial cells, bone, and glial cells and tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:209 as residues: Glu-15 to Thr-22, Glu-46 to Leu-62, Arg-103 to Glu-119, Gln-127 to Glu-132, Asn-152 to Trp-158, Gln-191 to Gln-210, Glu-264 to Thr-271, Tyr-282 to Leu-288, Trp-319 to Thr-331, Glu-335 to Ser-348, Ser-353 to Ser-358, Asp-382 to Asn-392.

The tissue distribution and homology to I-TRAF indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of inflammatory diseases, including rheumatoid arthritis, sepsis, inflammatory bowel disease, and psoriasis, particularly where tumor necrosis factor is known to be involved.

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FEATURES OF PROTEIN ENCODED BY GENE NO: 86

This gene has homology with a candidate gene involved in X-linked Retinopathy reported by Wong and colleagues (Genomics 15:467-471 (1993)).

This gene is expressed in a T-cell line.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, inflammation and autoimmune diseases; T-cell lymphoma. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the immune system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues and cell types (e.g., T-cells and other blood cells, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma, urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of inflammatory disorders such as sepsis, inflammatory bowel disease, psoriasis, and rheumatoid arthritis as well as autoimmune disease such as lupus. It could also be useful in immune modulation and in the process of immune surveillance. The present invention can be used diagnostically and therapeutically to treat X-linked Retinopathy.

25 FEATURES OF PROTEIN ENCODED BY GENE NO: 87

This gene is expressed in human brain tissue.

Therefore, polynucleotides and polypeptides of the invention are useful as reagents for differential identification of the tissue(s) or cell type(s) present in a biological sample and for diagnosis of diseases and conditions, which include, but are not limited to, brain disorders; neurodegenerative disorders; tumors of a brain origin. Similarly, polypeptides and antibodies directed to these polypeptides are useful in providing immunological probes for differential identification of the tissue(s) or cell type(s). For a number of disorders of the above tissues or cells, particularly of the central nervous system, expression of this gene at significantly higher or lower levels may be routinely detected in certain tissues (e.g., brain and other tissue of the nervous system, and cancerous and wounded tissues) or bodily fluids (e.g., serum, plasma,

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urine, synovial fluid or spinal fluid) or another tissue or cell sample taken from an individual having such a disorder, relative to the standard gene expression level, i.e., the expression level in healthy tissue or bodily fluid from an individual not having the disorder. Preferred epitopes include those comprising a sequence shown in SEQ ID NO:211 as residues: Cys-32 to Tyr-38.

The tissue distribution indicates that polynucleotides and polypeptides corresponding to this gene are useful for treatment and diagnosis of CNS disorders such as epilepsy, paranoia, depression, Alzheimer's disease, and schizophrenia. It could be useful in the survival and/or proliferation of neurons and could effect neuronal regeneration.

Last AA of ORF	30	44	69	- -	38	22	109
First AA of Secreted Portion		27	45	26	25	23	21
		26	44	25	24	22	20
First AA of Sig Pep			_	1	1	_	-
AA SEQ ID NO: Y	125	126	212	213	127	128	129
5' NT of First AA of Signal Pep	353	128	170	413	66	006	103
of of start	353	128	170	413	66	006	103
s, NT of Clone Seq.	1607	1786	1487	1637	1212	2061	733
Total Clone Clone NT Seq. Seq. Seq. Seq.	247	87	19	394		882	10
Total NT Seq.	1679	1830	1487	1653	1212	2061	1412
SEQ SEQ S.S.S.S.S.S.S.S.S.S.S.S.S.S.S.S.S.S.S.	Ξ	12	86	66	13	14	15
Vector	Uni-ZAP XR	Uni-ZAP XR	pBluescript	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR
ATCC Deposit Nr and Date	97923 03/07/97 209071	97923 03/07/97 209071	xxxxx 03/19/98	209641	97923 97923 03/07/97 209071	97923 03/07/97 209071	03/22/97 97923 03/07/97 209071 05/22/97
cDNA Clone ID	HAGEW82	HAGFY16	HBMCF37	HFLQB16	HALAA60	HAPBL78	HASAV70
Gene No.	-	2	2	2	3	4	ν

Last AA of ORF	62	29	52	56	215	48
First AA of Secreted Portion	18		24	81	61	27
Last AA of Sig Pep	17		23	17	8	26
First AA of Sig Pep		-	1	_		-
¥ŠEŽEŽ	130	131	132	133	134	135
S' NT First SEQ of AA For Start Signal NO: Start Codon Pep Y F	538	181	98	192	401	793
5' NT of Start Codon	538	181	98	192	401	793
3° NT of Clone Seq.	880	683	1007	1393	1070	2011
5' NT of Clone Seq.	276		98	132	277	614
FT S' NT3' NT S' Of S' O	1052	683	1054	1393	1215	2042
NT SEQ ID NO:	16	17	18	19	20	21
Vector	Uni-ZAP XR					
ATCC Deposit Nr and Date	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97
cDNA Clone ID	HBNAF22	HBNBL77	HCDDR90	нсеег50	HCEMU42	HCENE16
Gene No.	9	7	&	6	10	Ξ

Last AA of ORF	<i>L</i> 9	51	539	08	26	48	200
First AA of Secreted Portion	24	30	31	23	27	37	28
Last AA of Sig Pep	23	29	30	22	26	36	27
First AA of Sig Pep			_	_	_		_
AA SEQ ID NO: Y	136	137	138	214	139	215	140
5' NT of First AA of Signal Pep	69	89	808	515	961	295	70
5' NT of Start Codon	69	68	808	515	196	295	70
S' NT 3' NT of of Clone Clone Seq. Seq.	1872	289	3532	1115	907	734	717
S' NT of Clone (Seq.	21	1	2821	435	171	25	_
Total NT Seq.	1872	289	3533	1145	1148	734	717
NT SEQ ID NO:	22	23	24	100	25	101	26
Vector	Uni-ZAP XR	ZAP Express	Uni-ZAP XR	Uni-ZAP XR	pBluescript	Uni-ZAP XR	Uni-ZAP XR
ATCC Deposit Nr and Date	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	209179 07/24/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97
cDNA Clone ID	HMSJJ74	HCUBF15	HE2DE47	HE2DE47	HKMLH01	HE6DG34	HE9DG49
Gene No.	12	13	41	4	15	15	16

Last AA of ORF	202	215	185	101	=	19
First AA of Secreted Portion	29	23	26	43	31	
First Last AA AA of of of Sig Sig, Pep Pep	28	22	52.	42	30	
First AA of Sig Pep	_	-	1	1	1	-
AA SEQ BD: Y	216	141	217	142	143	144
of AA For SEQ AA of ID Signal NO: Pep Y	78	38	149	128	294	496
of of Start Sodon	78	38	149	128	294	496
3' NT of Clone Seq.	713	6601	0801	941	756	2093
Sop. Seq. Clone Seq. Clone Seq. Clone Clone Clone Clone Clone Clone Clone Clone Seq. Clone Seq. Clone Seq. Clone Seq. Clone Seq. Clone Clone Seq. Clone Seq. Clone Clone Seq. Clone Seq. Clone Seq. Clone Seq. Clone Clone Seq. Clone Seq. Clone Clone Seq. Clone Clone Clone Seq. Clone Clone Clone Seq. Clone Cl	17	_		171	62	408
Total NT Seq.	713	6601	1080	941	756	2100
XÖBÖX	102	27	103	28	A	30
Vector	Uni-ZAP XR					
ATCC Deposit Nr and Date	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97
cDNA Clone ID	HE9DG49	HELBA06	HELBA06	HSLFM29	HELBW38	HETHN28
Gene No.	16	17	17	81	19	20

Last AA of	ORF	67	8	_	38	130	31	13
First AA of Secreted		C	67		17	27	22	
Last of Of	Pep	C	87		91	26	21	
First AA of Sio	Pep		_		_			
SEQ AS	į 🖈	145	146	147	148	149		151
5' NT of AA I First SEQ AA of ID Signal NO:	Pep	267	21	210	242	178	144	1104
S' NT of	Codon	567	21	210	242	178	144	1104
3' NT of Clone	Sed.	1392	409	1322	710	1161	938	1581
S' NT 3' NT of of Clone Clone		475	-		_	110	_	974
Total	Seq.	1448	456	1326	710	1188	956	1603
SEQ	X S X	31	32	33	34	35	36	37
	Vector	Uni-ZAP XR	Uni-ZAP XR					
ATCC Deposit	Nr and Date	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/27/97	97923 03/07/97 209071 05/27/97	97923 03/07/97 209071 05/22/97	97923 03/07/97 209071 05/22/97	97924 03/07/97
	cDNA Clone ID	HFCDK17	HFEAF41	HFKFL13	HFSBG13	HFTBE43	HFTDJ36	HKTAC77
	Gene No.	21	22	23	24	25	26	27

Last AA of ORF	7	29	25	194	90	30	68	68	88	173	137	47	44
First AA of Secreted Portion		33		33	61	31	20	23	61	21	21	28	28
Last AA of Sig Pep		32		32	81	30	19	22	18	20	20	27	27
First AA of Sig Pep	1	1	1	I	1	1	-	_	1	I	1	_	-
AA SEQ ID NO: Y	152	153	154	551	156	157	158	218	159	160	219	220	161
5' NT of First AA of Signal Pep	209	119	581	126	43	171	55	58	17	15	72	54	269
5' NT of Start Codon		119	581	126	43	171	55	58	17	15	72	54	269
3' NT of Clone Seq.	1901	629	1793	1123	875	843	489	489	534	1374	640	1399	969
5' NT 3' NT of of Clone Seq. Seq.	55	1	408	13	_		3	9	1	1	58	40	-
Total NT Seq.	6801	629	1964	1522	875	843	489	489	534	1374	640	1529	596
NT SEQ D NO:	38	39	40	41	42	43	44	104	45	46	105	901	47
Vector	pBluescript	pBluescript	Lambda ZAP II	Uni-ZAP XR	Uni-ZAP XR	pSport1	Uni-ZAP XR						
ATCC Deposit Nr and Date	97924 03/07/97												
cDNA Clone ID	нгнзн36	96ASHTH	98ОВОТН	HLTBX31	HLTCJ63	HMKAH44	HMQAJ64	HMQAJ64	HOABG65	HODCL36	HODCL36	HODCL36	HODCL50
Gene No.	28	29	30	31	32	33	34	34	35	36	36	36	37

Last AA of ORF	22	69	322	69	319	82	30	71	280	42	22	326	183
First AA of Secreted Portion		18	20	32	61	22		19	31	31		20	24
Last AA of Sig Pep		17	19	31	60	21		18	30	30		19	23
First AA of Sig Pep	1		-		1	1	1	1	1	1	1	1	1
AA SEQ D NO: Y	162	163	164	221	165	222	166	167	168	223	169	170	224
5' NT of First AA of Signal Pep	170	638	99	928	150	239	432	142	25	433	217	23	35
5' NT of Start Codon	170	638	99	928	150	239	432	142	25	433	217	23	35
3' NT of Clone Seq.	822	2020	2432	2435	2340	791	109	337	1141	1166	1148	809	586
5' NT 3' NT of of Clone Clone Seq.	66	569	848	849	1627	92	188		1	21	63	164	4
Total NT Seq.	851	2020	2432	2435	2340	805	601	359	1141	9911	1560	1507	586
NT SEQ ID NO:	48	46	50	107	51	801	52	53	54	109	55	99	110
Vector	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	pBluescript SK-	pBluescript SK-	Uni-ZAP XR						
ATCC Deposit Nr and Date	97924 03/07/97	97924 03/07/97	97924 03/07/97	97924 03/07/97	97924 03/07/97	97924 03/07/97	97924 03/07/97	97924 03/07/97	97924 03/07/97	97924 03/07/97	97924 03/07/97	97924 03/07/97	97924 03/07/97
cDNA Clone ID	HODCV74	HODCZ16	HTOEU03	HTOEU03	HPBCJ74	HPBCJ74	HPMBU33	HSAUL66	HSIDQ18	HSIDQ18	HSJBB37	HSJBQ79	HSJBQ79
Gene No.	38	39	40	40	41	41	42	43	44	44	45	46	46

Last AA of ORF	89	158	70	122	128	6	371
First AA of Secreted Portion	36	16	20	19	31		2
Last AA of Sig Pep	35	15	61	81	30		_
First AA of Sig Pep	_	_	1	_	_	_	
AA SEQ ID NO: Y	171	172	225	173	174	226	175
5' NT of First AA of Signal Pep	83	163	155	115	52	829	114
5' NT of Start Codon	83	163	155	115	52	829	114
3' NT of Clone Seq.	450	1147	1134	LLL	865	1333	1554
5' NT 3' NT of of Clone Seq. Seq.	_				48	594	443
Total NT Seq.	450	1147	1134	777	1191	1333	1580
NT SEQ NO:	57	28		59	09	112	19
Vector	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	pBluescript	pBluescript	Uni-ZAP XR
ATCC Deposit Nr and Date	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	209235 09/04/97
cDNA Clone ID	HTEGA76	HTEJN13	HTEJN13	HTHBL86	HTSF071	HTSF071	HAPNO80
Gene No.	47	48	48	49	50	50	51

Last AA of ORF	137	215	54	22	102	47
First AA of Secreted Portion	29	29	33	21	34	39
	28	28	32	20	33	38
First Last AA AA of of Sig Sig Pep	1		_	1	.	_
AA SEQ ID NO: Y	227	176	177	178	179	180
5' NT of First AA of Signal Pep	244	. 182	76	150	231	703
of of Start	244	182	97	150	231	703
3' NT of Clone Seq.	708	1034	361	1638	1303	1011
S' NT 3' NT 5 of of 5 of S of Clone Clone NT Seq. Seq.	249	105			35	655
Total NT Seq.	1015	1117	361	1668	1353	1011
X SEQ	113	79	63	64	92	99
Vector	Uni-ZAP XR	pBluescript	pSport1	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR
ATCC Deposit Nr and Date	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97
cDNA Clone ID	HAUCC47	HBMCL41	HCFLD84	HE8EM69	HE8EZ48	HEBGF73
Gene No.	51	52	53	54	55	56

Last AA of ORF	95	94	26	01	64	21
First AA of Secreted Portion	36	30	22		20	22
Last AA of Sig Pep	35	29	21		61	21
First AA of Sig Pep		_	-	-		_
¥SEQ YÖ: BÖ	181	182	183	184	185	186
5' NT of First AA of Signal Pep	459	63	839	270	272	127
5' NT of Start Codon	459	63	839	270	272	127
3' NT of Clone Seq.	1090	995	1581	711	935	484
S' NT 3' NT of of Clone Seq.	267	-	765	∞		113
Tota NT Seq	1193	260	1657	711	935	504
SEQ NA	<i>L</i> 9	89	69	70	71	72
Vector	Uni-ZAP XR	Uni-ZAP XR	pBluescript	Lambda ZAP II	Lambda ZAP II	Lambda ZAP II
ATCC Deposit Nr and Date	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97
cDNA Clone ID	HFEBF41	HFRBU14	HFVGZ79	нн <u>дсм</u> 76	HHGCO88	HHGCP52
Gene No.	57	28	59	09	19	62

Last AA of ORF	131	89	44	49	22	169
First AA of Secreted Portion	61	33	28	37	12	15
	18	32	27	36	11	14
First Last AA AA of of Sig Sig Pep Pep		-		1		
AA SEQ ID NO: Y	187	188	189	190	228	192
of AA Fi of AA of D of AA of D o Start Signal NO: S	96	248	630	191	575	187
	96	248	630	167		187
3' NT of Clone Seq.	620	581	1786	800	1076	1888
S' NT 3' NT of of Clone Clone Seq. Seq.	-	951	537	116	398	18
Total NT Seq.	620	581	1843	1441	1076	2776
SEQ NÖ:	73	74	75	9/	114	78
Vector	Lambda ZAP II	Lambda ZAP II	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	pCMVSport 3.0
ATCC Deposit Nr and Date	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97	97958 03/13/97 209072 05/22/97
cDNA Clone ID	HHGDB72	HHGDI71	HHSD145	HHSEB66	HJPAZ83	HLDBO49
Gene No.	63	64	65	99	19	89

Last AA of ORF	65	131	91	175	69	24	72
First AA of Secreted Portion	23	23	33	24	27	21	26
Last AA of Sig Pep	22	22	32	23	26	20	25
First AA of Sig Pep	-	1		-			-
AA SEQ D NÖ:	193	229	194	195	961	197	861
of AA First SEQ AA of ID Signal NO: Pep Y	534	534	40	238	286	28	14
5' NT of Start Codon	534	234	40	238	286	58	4
S' NT 3' NT of Olone Clone Seq. Seq.	1480	1487	1077	780	770	481	623
S' NT 3' NT of of Clone Clone Seq.	401	401	33	<u>&</u>	101	_	_
Total NT Seq.	1525	1487	1563	1020	770	481	644
SEQ SEQ NO:	62	115	08	8	82	83	84
Vector	pCMVSport 3.0	pCMVSport 3.0	Uni-ZAP XR	Uni-Zap XR	Uni-ZAP XR	Uni-ZAP XR	pCMVSport 3.0
ATCC Deposit Nr and Date	97958 03/13/97 209072 05/22/97	209226 08/28/97	97958 03/13/97 209072 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97
cDNA Clone ID	HLDBQ19	HLDBQ19	HMSGT42	HMWIC78	HTTCT79	HNGJU84	HNTAC73
Gene No.	69	69	70	71	72	73	74

	288	27	623	09	648	28
First AA of Secreted Portion	13		31	33	31	22
Last AA of Sig Pep	12		30	32	30	21
First Last AA AA of of Sig Sig Pep Pep		_		1		_
AA SEQ ID NO: Y	199	230	200	231	201	232
of AA For SEQ AA of ID Signal NO:	86	545	99	477	251	677
	86		56	477	251	677
3' NT of Clone Seq.	1284	1283	1747	1747	2566	1098
NT SEQ of of 5' N D Total Clone Clone of Stan Stan Seq. Seq. Seq. Cod	435	428	290	288	1843 2566	375
Total NT Seq.	1351	1350	2527	2527	2566	8601
SEQ NÖ:	85	116	98	117	87	118
Vector	Uni-ZAP XR					
ATCC Deposit Nr and Date	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97
cDNA Clone ID	HOSE145	HOSE145	HOSFD58	HOSFD58	HSAUM95	HSAUM95
Gene No.	75	75	76	76	77	11

Last AA of ORF	54	265	17	314	206	194
First AA of Secreted Portion	33	12		20	21	70
Last AA of of Sig Pep	32	Ξ		61	20	69
First AA of Sig Pep	—		_	I	-	_
AA SEQ NÖ:	202	203	233	204	205	206
of AA For SEQ AA of DA Signal NO: Pep Y H	83	881	315	92	414	157
of of Start Sodon	83	188	315	92	414	157
5' NT 3' NT of of Clone Clone Seq. Seq.	540	1165	1166	2449	2058	1411
S' NT 3' NT of of Soft Clone NT Seq. Seq.		152	152	1149	476	345
Total NT Seq.	540	1863	1679	2478	2058	1411
NT SEQ D NO:	88	68	611	06	16	92
Vector	Uni-ZAP XR	pBluescript				
ATCC Deposit Nr and Date	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97
cDNA Clone ID	HSAUR67	HSKDI81	HSKD181	HSKDW91	HTLEX50	HSKHL65
Gene No.	78	79	79	80	8	82

	7.1	329	95	57	391	25
First AA of Secreted Portion	38	31	20	21	2	22
	37	30	61	20	_	21
irst AA of Sig	,					_
AA SEQ ID NO: Y	235	207	236	208	209	210
of AA F of Example of AA of AA of D Signal NO: Signal NO: Pep Y I	526	397	228	445	523	117
S' NT of Start	526	397	228	445	523	117
3' NT of Clone Seq.	1411	2184	2063	809	2394	672
S' NT 3' NT of of School Clone Seq. Seq.	345	147	138	524	481	_
Total NT Seq.	1411	2187	2256	757	2394	672
SEQ NO:	121	93	122	94	95	96
Vector	pBluescript	Uni-ZAP XR	Uni-ZAP XR	Uni-ZAP XR	ZAP Express	Uni-ZAP XR
ATCC Deposit Nr and Date	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97	97957 03/13/97 209073 05/22/97
cDNA Clone ID	HSKHL65	HHFGA11	HHFGAII	HWTBL40	HBXFG80	HCACY32
Gene No.	82	83	83	84	85	98

Last AA of ORF	37
S' NTofAAFirstLastT FirstSEQAAAAFirst AALastAA ofIDofofAAtSignalNO:SigSigSecretedofnPepYPepPortionORF	21
Last AA of Sig Pep	20
First AA of Sig Pep	211 1
AA SEQ D NO: Y	211
5' NT of First AA of Signal Pep	207
S' NT 3' NT of of S' NT I of of Clone Clone of A Start	207
3' NT of Clone Seq.	1419
5' NT of Clone Seq.	1
Tot N	1419
SEQ NO:	97
Vector	97957 Uni-ZAP XR 97 1419 1 1419 3/13/97 209073 5/22/97
ATCC Deposit Nr and Date	97957 03/13/97 209073 05/22/97
cDNA Clone ID	HCED021
Gene No.	Ĺď

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Table 1 summarizes the information corresponding to each "Gene No." described above. The nucleotide sequence identified as "NT SEQ ID NO:X" was assembled from partially homologous ("overlapping") sequences obtained from the "cDNA clone ID" identified in Table 1 and, in some cases, from additional related DNA clones. The overlapping sequences were assembled into a single contiguous sequence of high redundancy (usually three to five overlapping sequences at each nucleotide position), resulting in a final sequence identified as SEQ ID NO:X.

The cDNA Clone ID was deposited on the date and given the corresponding deposit number listed in "ATCC Deposit No:Z and Date." Some of the deposits contain multiple different clones corresponding to the same gene. "Vector" refers to the type of vector contained in the cDNA Clone ID.

"Total NT Seq." refers to the total number of nucleotides in the contig identified by "Gene No." The deposited clone may contain all or most of these sequences, reflected by the nucleotide position indicated as "5' NT of Clone Seq." and the "3' NT of Clone Seq." of SEQ ID NO:X. The nucleotide position of SEQ ID NO:X of the putative start codon (methionine) is identified as "5' NT of Start Codon." Similarly, the nucleotide position of SEQ ID NO:X of the predicted signal sequence is identified as "5' NT of First AA of Signal Pep."

The translated amino acid sequence, beginning with the methionine, is identified as "AA SEQ ID NO:Y," although other reading frames can also be easily translated using known molecular biology techniques. The polypeptides produced by these alternative open reading frames are specifically contemplated by the present invention.

The first and last amino acid position of SEQ ID NO:Y of the predicted signal peptide is identified as "First AA of Sig Pep" and "Last AA of Sig Pep." The predicted first amino acid position of SEQ ID NO:Y of the secreted portion is identified as "Predicted First AA of Secreted Portion." Finally, the amino acid position of SEQ ID NO:Y of the last amino acid in the open reading frame is identified as "Last AA of ORF."

SEQ ID NO:X and the translated SEQ ID NO:Y are sufficiently accurate and otherwise suitable for a variety of uses well known in the art and described further below. For instance, SEQ ID NO:X is useful for designing nucleic acid hybridization probes that will detect nucleic acid sequences contained in SEQ ID NO:X or the cDNA contained in the deposited clone. These probes will also hybridize to nucleic acid molecules in biological samples, thereby enabling a variety of forensic and diagnostic methods of the invention. Similarly, polypeptides identified from SEQ ID NO:Y may be used to generate antibodies which bind specifically to the secreted proteins encoded by the cDNA clones identified in Table 1.

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Nevertheless, DNA sequences generated by sequencing reactions can contain sequencing errors. The errors exist as misidentified nucleotides, or as insertions or deletions of nucleotides in the generated DNA sequence. The erroneously inserted or deleted nucleotides cause frame shifts in the reading frames of the predicted amino acid sequence. In these cases, the predicted amino acid sequence diverges from the actual amino acid sequence, even though the generated DNA sequence may be greater than 99.9% identical to the actual DNA sequence (for example, one base insertion or deletion in an open reading frame of over 1000 bases).

Accordingly, for those applications requiring precision in the nucleotide sequence or the amino acid sequence, the present invention provides not only the generated nucleotide sequence identified as SEQ ID NO:X and the predicted translated amino acid sequence identified as SEQ ID NO:Y, but also a sample of plasmid DNA containing a human cDNA of the invention deposited with the ATCC, as set forth in Table 1. The nucleotide sequence of each deposited clone can readily be determined by sequencing the deposited clone in accordance with known methods. The predicted amino acid sequence can then be verified from such deposits. Moreover, the amino acid sequence of the protein encoded by a particular clone can also be directly determined by peptide sequencing or by expressing the protein in a suitable host cell containing the deposited human cDNA, collecting the protein, and determining its sequence.

The present invention also relates to the genes corresponding to SEQ ID NO:X, SEQ ID NO:Y, or the deposited clone. The corresponding gene can be isolated in accordance with known methods using the sequence information disclosed herein. Such methods include preparing probes or primers from the disclosed sequence and identifying or amplifying the corresponding gene from appropriate sources of genomic material.

Also provided in the present invention are species homologs. Species homologs may be isolated and identified by making suitable probes or primers from the sequences provided herein and screening a suitable nucleic acid source for the desired homologue.

The polypeptides of the invention can be prepared in any suitable manner. Such polypeptides include isolated naturally occurring polypeptides, recombinantly produced polypeptides, synthetically produced polypeptides, or polypeptides produced by a combination of these methods. Means for preparing such polypeptides are well understood in the art.

The polypeptides may be in the form of the secreted protein, including the mature form, or may be a part of a larger protein, such as a fusion protein (see below).

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It is often advantageous to include an additional amino acid sequence which contains secretory or leader sequences, pro-sequences, sequences which aid in purification, such as multiple histidine residues, or an additional sequence for stability during recombinant production.

The polypeptides of the present invention are preferably provided in an isolated form, and preferably are substantially purified. A recombinantly produced version of a polypeptide, including the secreted polypeptide, can be substantially purified by the one-step method described in Smith and Johnson, Gene 67:31-40 (1988). Polypeptides of the invention also can be purified from natural or recombinant sources using antibodies of the invention raised against the secreted protein in methods which are well known in the art.

Signal Sequences

Methods for predicting whether a protein has a signal sequence, as well as the cleavage point for that sequence, are available. For instance, the method of McGeoch, Virus Res. 3:271-286 (1985), uses the information from a short N-terminal charged region and a subsequent uncharged region of the complete (uncleaved) protein. The method of von Heinje, Nucleic Acids Res. 14:4683-4690 (1986) uses the information from the residues surrounding the cleavage site, typically residues -13 to +2, where +1 indicates the amino terminus of the secreted protein. The accuracy of predicting the cleavage points of known mammalian secretory proteins for each of these methods is in the range of 75-80%. (von Heinje, supra.) However, the two methods do not always produce the same predicted cleavage point(s) for a given protein.

In the present case, the deduced amino acid sequence of the secreted polypeptide was analyzed by a computer program called SignalP (Henrik Nielsen et al., Protein Engineering 10:1-6 (1997)), which predicts the cellular location of a protein based on the amino acid sequence. As part of this computational prediction of localization, the methods of McGeoch and von Heinje are incorporated. The analysis of the amino acid sequences of the secreted proteins described herein by this program provided the results shown in Table 1.

As one of ordinary skill would appreciate, however, cleavage sites sometimes vary from organism to organism and cannot be predicted with absolute certainty. Accordingly, the present invention provides secreted polypeptides having a sequence shown in SEQ ID NO:Y which have an N-terminus beginning within 5 residues (i.e., + or - 5 residues) of the predicted cleavage point. Similarly, it is also recognized that in some cases, cleavage of the signal sequence from a secreted protein is not entirely

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uniform, resulting in more than one secreted species. These polypeptides, and the polynucleotides encoding such polypeptides, are contemplated by the present invention.

Moreover, the signal sequence identified by the above analysis may not necessarily predict the naturally occurring signal sequence. For example, the naturally occurring signal sequence may be further upstream from the predicted signal sequence. However, it is likely that the predicted signal sequence will be capable of directing the secreted protein to the ER. These polypeptides, and the polynucleotides encoding such polypeptides, are contemplated by the present invention.

Polynucleotide and Polypeptide Variants

"Variant" refers to a polynucleotide or polypeptide differing from the polynucleotide or polypeptide of the present invention, but retaining essential properties thereof. Generally, variants are overall closely similar, and, in many regions, identical to the polynucleotide or polypeptide of the present invention.

By a polynucleotide having a nucleotide sequence at least, for example, 95% "identical" to a reference nucleotide sequence of the present invention, it is intended that the nucleotide sequence of the polynucleotide is identical to the reference sequence except that the polynucleotide sequence may include up to five point mutations per each 100 nucleotides of the reference nucleotide sequence encoding the polypeptide. In other words, to obtain a polynucleotide having a nucleotide sequence at least 95% identical to a reference nucleotide sequence, up to 5% of the nucleotides in the reference sequence may be deleted or substituted with another nucleotide, or a number of nucleotides up to 5% of the total nucleotides in the reference sequence may be inserted into the reference sequence. The query sequence may be an entire sequence shown in Table 1, the ORF (open reading frame), or any fragement specified as described herein.

As a practical matter, whether any particular nucleic acid molecule or polypeptide is at least 90%, 95%, 96%, 97%, 98% or 99% identical to a nucleotide sequence of the presence invention can be determined conventionally using known computer programs. A preferred method for determing the best overall match between a query sequence (a sequence of the present invention) and a subject sequence, also referred to as a global sequence alignment, can be determined using the FASTDB computer program based on the algorithm of Brutlag et al. (Comp. App. Biosci. (1990) 6:237-245). In a sequence alignment the query and subject sequences are both DNA sequences. An RNA sequence can be compared by converting U's to T's. The result of said global sequence alignment is in percent identity. Preferred parameters used in a FASTDB alignment of DNA sequences to calculate percent identity are:

Matrix=Unitary, k-tuple=4, Mismatch Penalty=1, Joining Penalty=30, Randomization

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Group Length=0, Cutoff Score=1, Gap Penalty=5, Gap Size Penalty 0.05, Window Size=500 or the length of the subject nucleotide sequence, whichever is shorter.

If the subject sequence is shorter than the query sequence because of 5' or 3' deletions, not because of internal deletions, a manual correction must be made to the results. This is because the FASTDB program does not account for 5' and 3' truncations of the subject sequence when calculating percent identity. For subject sequences truncated at the 5' or 3' ends, relative to the the query sequence, the percent identity is corrected by calculating the number of bases of the query sequence that are 5' and 3' of the subject sequence, which are not matched/aligned, as a percent of the total bases of the query sequence. Whether a nucleotide is matched/aligned is determined by results of the FASTDB sequence alignment. This percentage is then subtracted from the percent identity, calculated by the above FASTDB program using the specified parameters, to arrive at a final percent identity score. This corrected score is what is used for the purposes of the present invention. Only bases outside the 5' and 3' bases of the subject sequence, as displayed by the FASTDB alignment, which are not matched/aligned with the query sequence, are calculated for the purposes of manually adjusting the percent identity score.

For example, a 90 base subject sequence is aligned to a 100 base query sequence to determine percent identity. The deletions occur at the 5' end of the subject sequence and therefore, the FASTDB alignment does not show a matched/alignement of the first 10 bases at 5' end. The 10 unpaired bases represent 10% of the sequence (number of bases at the 5' and 3' ends not matched/total number of bases in the query sequence) so 10% is subtracted from the percent identity score calculated by the FASTDB program. If the remaining 90 bases were perfectly matched the final percent identity would be 90%. In another example, a 90 base subject sequence is compared with a 100 base query sequence. This time the deletions are internal deletions so that there are no bases on the 5' or 3' of the subject sequence which are not matched/aligned with the query. In this case the percent identity calculated by FASTDB is not manually corrected. Once again, only bases 5' and 3' of the subject sequence which are not matched/aligned with the query sequence are manually corrected for. No other manual corrections are to made for the purposes of the present invention.

By a polypeptide having an amino acid sequence at least, for example, 95% "identical" to a query amino acid sequence of the present invention, it is intended that the amino acid sequence of the subject polypeptide is identical to the query sequence except that the subject polypeptide sequence may include up to five amino acid alterations per each 100 amino acids of the query amino acid sequence. In other words, to obtain a polypeptide having an amino acid sequence at least 95% identical to a query

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amino acid sequence, up to 5% of the amino acid residues in the subject sequence may be inserted, deleted, (indels) or substituted with another amino acid. These alterations of the reference sequence may occur at the amino or carboxy terminal positions of the reference amino acid sequence or anywhere between those terminal positions, interspersed either individually among residues in the reference sequence or in one or more contiguous groups within the reference sequence.

As a practical matter, whether any particular polypeptide is at least 90%, 95%, 96%, 97%, 98% or 99% identical to, for instance, the amino acid sequences shown in Table 1 or to the amino acid sequence encoded by deposited DNA clone can be determined conventionally using known computer programs. A preferred method for determing the best overall match between a query sequence (a sequence of the present invention) and a subject sequence, also referred to as a global sequence alignment, can be determined using the FASTDB computer program based on the algorithm of Brutlag et al. (Comp. App. Biosci. (1990) 6:237-245). In a sequence alignment the query and subject sequences are either both nucleotide sequences or both amino acid sequences. The result of said global sequence alignment is in percent identity. Preferred parameters used in a FASTDB amino acid alignment are: Matrix=PAM 0, k-tuple=2, Mismatch Penalty=1, Joining Penalty=20, Randomization Group Length=0, Cutoff Score=1, Window Size=sequence length, Gap Penalty=5, Gap Size Penalty=0.05, Window Size=500 or the length of the subject amino acid sequence, whichever is shorter.

If the subject sequence is shorter than the query sequence due to N- or Cterminal deletions, not because of internal deletions, a manual correction must be made to the results. This is because the FASTDB program does not account for N- and Cterminal truncations of the subject sequence when calculating global percent identity. For subject sequences truncated at the N- and C-termini, relative to the the query sequence, the percent identity is corrected by calculating the number of residues of the query sequence that are N- and C-terminal of the subject sequence, which are not matched/aligned with a corresponding subject residue, as a percent of the total bases of the query sequence. Whether a residue is matched/aligned is determined by results of the FASTDB sequence alignment. This percentage is then subtracted from the percent identity, calculated by the above FASTDB program using the specified parameters, to arrive at a final percent identity score. This final percent identity score is what is used for the purposes of the present invention. Only residues to the N- and C-termini of the subject sequence, which are not matched/aligned with the query sequence, are considered for the purposes of manually adjusting the percent identity score. That is, only query residue positions outside the farthest N- and C-terminal residues of the subject sequence.

For example, a 90 amino acid residue subject sequence is aligned with a 100 residue query sequence to determine percent identity. The deletion occurs at the Nterminus of the subject sequence and therefore, the FASTDB alignment does not show a matching/alignment of the first 10 residues at the N-terminus. The 10 unpaired residues represent 10% of the sequence (number of residues at the N- and C- termini 5 not matched/total number of residues in the query sequence) so 10% is subtracted from the percent identity score calculated by the FASTDB program. If the remaining 90 residues were perfectly matched the final percent identity would be 90%. In another example, a 90 residue subject sequence is compared with a 100 residue query sequence. This time the deletions are internal deletions so there are no residues at the N- or C-10 termini of the subject sequence which are not matched/aligned with the query. In this case the percent identity calculated by FASTDB is not manually corrected. Once again, only residue positions outside the N- and C-terminal ends of the subject sequence, as displayed in the FASTDB alignment, which are not matched/aligned with the query sequnce are manually corrected for. No other manual corrections are to made for the 15 purposes of the present invention.

The variants may contain alterations in the coding regions, non-coding regions, or both. Especially preferred are polynucleotide variants containing alterations which produce silent substitutions, additions, or deletions, but do not alter the properties or activities of the encoded polypeptide. Nucleotide variants produced by silent substitutions due to the degeneracy of the genetic code are preferred. Moreover, variants in which 5-10, 1-5, or 1-2 amino acids are substituted, deleted, or added in any combination are also preferred. Polynucleotide variants can be produced for a variety of reasons, e.g., to optimize codon expression for a particular host (change codons in the human mRNA to those preferred by a bacterial host such as E. coli).

Naturally occurring variants are called "allelic variants," and refer to one of several alternate forms of a gene occupying a given locus on a chromosome of an organism. (Genes II, Lewin, B., ed., John Wiley & Sons, New York (1985).) These allelic variants can vary at either the polynucleotide and/or polypeptide level. Alternatively, non-naturally occurring variants may be produced by mutagenesis techniques or by direct synthesis.

Using known methods of protein engineering and recombinant DNA technology, variants may be generated to improve or alter the characteristics of the polypeptides of the present invention. For instance, one or more amino acids can be deleted from the N-terminus or C-terminus of the secreted protein without substantial loss of biological function. The authors of Ron et al., J. Biol. Chem. 268: 2984-2988 (1993), reported variant KGF proteins having heparin binding activity even after

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deleting 3, 8, or 27 amino-terminal amino acid residues. Similarly, Interferon gamma exhibited up to ten times higher activity after deleting 8-10 amino acid residues from the carboxy terminus of this protein. (Dobeli et al., J. Biotechnology 7:199-216 (1988).)

Moreover, ample evidence demonstrates that variants often retain a biological activity similar to that of the naturally occurring protein. For example, Gayle and coworkers (J. Biol. Chem 268:22105-22111 (1993)) conducted extensive mutational analysis of human cytokine IL-1a. They used random mutagenesis to generate over 3,500 individual IL-1a mutants that averaged 2.5 amino acid changes per variant over the entire length of the molecule. Multiple mutations were examined at every possible amino acid position. The investigators found that "[m]ost of the molecule could be altered with little effect on either [binding or biological activity]." (See, Abstract.) In fact, only 23 unique amino acid sequences, out of more than 3,500 nucleotide sequences examined, produced a protein that significantly differed in activity from wild-type.

Furthermore, even if deleting one or more amino acids from the N-terminus or C-terminus of a polypeptide results in modification or loss of one or more biological functions, other biological activities may still be retained. For example, the ability of a deletion variant to induce and/or to bind antibodies which recognize the secreted form will likely be retained when less than the majority of the residues of the secreted form are removed from the N-terminus or C-terminus. Whether a particular polypeptide lacking N- or C-terminal residues of a protein retains such immunogenic activities can readily be determined by routine methods described herein and otherwise known in the art.

Thus, the invention further includes polypeptide variants which show substantial biological activity. Such variants include deletions, insertions, inversions, repeats, and substitutions selected according to general rules known in the art so as have little effect on activity. For example, guidance concerning how to make phenotypically silent amino acid substitutions is provided in Bowie, J. U. et al., Science 247:1306-1310 (1990), wherein the authors indicate that there are two main strategies for studying the tolerance of an amino acid sequence to change.

The first strategy exploits the tolerance of amino acid substitutions by natural selection during the process of evolution. By comparing amino acid sequences in different species, conserved amino acids can be identified. These conserved amino acids are likely important for protein function. In contrast, the amino acid positions where substitutions have been tolerated by natural selection indicates that these positions are not critical for protein function. Thus, positions tolerating amino acid substitution could be modified while still maintaining biological activity of the protein.

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The second strategy uses genetic engineering to introduce amino acid changes at specific positions of a cloned gene to identify regions critical for protein function. For example, site directed mutagenesis or alanine-scanning mutagenesis (introduction of single alanine mutations at every residue in the molecule) can be used. (Cunningham and Wells, Science 244:1081-1085 (1989).) The resulting mutant molecules can then be tested for biological activity.

As the authors state, these two strategies have revealed that proteins are surprisingly tolerant of amino acid substitutions. The authors further indicate which amino acid changes are likely to be permissive at certain amino acid positions in the protein. For example, most buried (within the tertiary structure of the protein) amino acid residues require nonpolar side chains, whereas few features of surface side chains are generally conserved. Moreover, tolerated conservative amino acid substitutions involve replacement of the aliphatic or hydrophobic amino acids Ala, Val, Leu and Ile; replacement of the hydroxyl residues Ser and Thr; replacement of the acidic residues Asp and Glu; replacement of the amide residues Asn and Gln, replacement of the basic residues Lys, Arg, and His; replacement of the aromatic residues Phe, Tyr, and Trp, and replacement of the small-sized amino acids Ala, Ser, Thr, Met, and Gly.

Besides conservative amino acid substitution, variants of the present invention include (i) substitutions with one or more of the non-conserved amino acid residues, where the substituted amino acid residues may or may not be one encoded by the genetic code, or (ii) substitution with one or more of amino acid residues having a substituent group, or (iii) fusion of the mature polypeptide with another compound, such as a compound to increase the stability and/or solubility of the polypeptide (for example, polyethylene glycol), or (iv) fusion of the polypeptide with additional amino acids, such as an IgG Fc fusion region peptide, or leader or secretory sequence, or a sequence facilitating purification. Such variant polypeptides are deemed to be within the scope of those skilled in the art from the teachings herein.

For example, polypeptide variants containing amino acid substitutions of charged amino acids with other charged or neutral amino acids may produce proteins with improved characteristics, such as less aggregation. Aggregation of pharmaceutical formulations both reduces activity and increases clearance due to the aggregate's immunogenic activity. (Pinckard et al., Clin. Exp. Immunol. 2:331-340 (1967); Robbins et al., Diabetes 36: 838-845 (1987); Cleland et al., Crit. Rev. Therapeutic Drug Carrier Systems 10:307-377 (1993).)

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Polynucleotide and Polypeptide Fragments

In the present invention, a "polynucleotide fragment" refers to a short polynucleotide having a nucleic acid sequence contained in the deposited clone or shown in SEQ ID NO:X. The short nucleotide fragments are preferably at least about 15 nt, and more preferably at least about 20 nt, still more preferably at least about 30 nt, and even more preferably, at least about 40 nt in length. A fragment "at least 20 nt in length," for example, is intended to include 20 or more contiguous bases from the cDNA sequence contained in the deposited clone or the nucleotide sequence shown in SEQ ID NO:X. These nucleotide fragments are useful as diagnostic probes and primers as discussed herein. Of course, larger fragments (e.g., 50, 150, 500, 600, 2000 nucleotides) are preferred.

Moreover, representative examples of polynucleotide fragments of the invention, include, for example, fragments having a sequence from about nucleotide number 1-50, 51-100, 101-150, 151-200, 201-250, 251-300, 301-350, 351-400, 401-450, 451-500, 501-550, 551-600, 651-700, or 701 to the end of SEQ ID NO:X or the cDNA contained in the deposited clone. In this context "about" includes the particularly recited ranges, larger or smaller by several (5, 4, 3, 2, or 1) nucleotides, at either terminus or at both termini. Preferably, these fragments encode a polypeptide which has biological activity.

In the present invention, a "polypeptide fragment" refers to a short amino acid sequence contained in SEQ ID NO:Y or encoded by the cDNA contained in the deposited clone. Protein fragments may be "free-standing," or comprised within a larger polypeptide of which the fragment forms a part or region, most preferably as a single continuous region. Representative examples of polypeptide fragments of the invention, include, for example, fragments from about amino acid number 1-20, 21-40, 41-60, 61-80, 81-100, 102-120, 121-140, 141-160, or 161 to the end of the coding region. Moreover, polypeptide fragments can be about 20, 30, 40, 50, 60, 70, 80, 90, 100, 110, 120, 130, 140, or 150 amino acids in length. In this context "about" includes the particularly recited ranges, larger or smaller by several (5, 4, 3, 2, or 1) amino acids, at either extreme or at both extremes.

Preferred polypeptide fragments include the secreted protein as well as the mature form. Further preferred polypeptide fragments include the secreted protein or the mature form having a continuous series of deleted residues from the amino or the carboxy terminus, or both. For example, any number of amino acids, ranging from 1-60, can be deleted from the amino terminus of either the secreted polypeptide or the mature form. Similarly, any number of amino acids, ranging from 1-30, can be deleted from the carboxy terminus of the secreted protein or mature form. Furthermore, any

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combination of the above amino and carboxy terminus deletions are preferred. Similarly, polynucleotide fragments encoding these polypeptide fragments are also preferred.

Also preferred are polypeptide and polynucleotide fragments characterized by structural or functional domains, such as fragments that comprise alpha-helix and alpha-helix forming regions, beta-sheet and beta-sheet-forming regions, turn and turn-forming regions, coil and coil-forming regions, hydrophilic regions, hydrophobic regions, alpha amphipathic regions, beta amphipathic regions, flexible regions, surface-forming regions, substrate binding region, and high antigenic index regions.

Polypeptide fragments of SEQ ID NO:Y falling within conserved domains are specifically contemplated by the present invention. Moreover, polynucleotide fragments encoding these domains are also contemplated.

Other preferred fragments are biologically active fragments. Biologically active fragments are those exhibiting activity similar, but not necessarily identical, to an activity of the polypeptide of the present invention. The biological activity of the fragments may include an improved desired activity, or a decreased undesirable activity.

Epitopes & Antibodies

In the present invention, "epitopes" refer to polypeptide fragments having antigenic or immunogenic activity in an animal, especially in a human. A preferred embodiment of the present invention relates to a polypeptide fragment comprising an epitope, as well as the polynucleotide encoding this fragment. A region of a protein molecule to which an antibody can bind is defined as an "antigenic epitope." In contrast, an "immunogenic epitope" is defined as a part of a protein that elicits an antibody response. (See, for instance, Geysen et al., Proc. Natl. Acad. Sci. USA 81:3998- 4002 (1983).)

Fragments which function as epitopes may be produced by any conventional means. (See, e.g., Houghten, R. A., Proc. Natl. Acad. Sci. USA 82:5131-5135 (1985) further described in U.S. Patent No. 4,631,211.)

In the present invention, antigenic epitopes preferably contain a sequence of at least seven, more preferably at least nine, and most preferably between about 15 to about 30 amino acids. Antigenic epitopes are useful to raise antibodies, including monoclonal antibodies, that specifically bind the epitope. (See, for instance, Wilson et al., Cell 37:767-778 (1984); Sutcliffe, J. G. et al., Science 219:660-666 (1983).)

Similarly, immunogenic epitopes can be used to induce antibodies according to methods well known in the art. (See, for instance, Sutcliffe et al., supra; Wilson et al., supra; Chow, M. et al., Proc. Natl. Acad. Sci. USA 82:910-914; and Bittle, F. J. et

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al., J. Gen. Virol. 66:2347-2354 (1985).) A preferred immunogenic epitope includes the secreted protein. The immunogenic epitopes may be presented together with a carrier protein, such as an albumin, to an animal system (such as rabbit or mouse) or, if it is long enough (at least about 25 amino acids), without a carrier. However, immunogenic epitopes comprising as few as 8 to 10 amino acids have been shown to be sufficient to raise antibodies capable of binding to, at the very least, linear epitopes in a denatured polypeptide (e.g., in Western blotting.)

As used herein, the term "antibody" (Ab) or "monoclonal antibody" (Mab) is meant to include intact molecules as well as antibody fragments (such as, for example, Fab and F(ab')2 fragments) which are capable of specifically binding to protein. Fab and F(ab')2 fragments lack the Fc fragment of intact antibody, clear more rapidly from the circulation, and may have less non-specific tissue binding than an intact antibody. (Wahl et al., J. Nucl. Med. 24:316-325 (1983).) Thus, these fragments are preferred. as well as the products of a FAB or other immunoglobulin expression library. Moreover, antibodies of the present invention include chimeric, single chain, and

humanized antibodies.

Fusion Proteins

Any polypeptide of the present invention can be used to generate fusion. proteins. For example, the polypeptide of the present invention, when fused to a second protein, can be used as an antigenic tag. Antibodies raised against the polypeptide of the present invention can be used to indirectly detect the second protein by binding to the polypeptide. Moreover, because secreted proteins target cellular locations based on trafficking signals, the polypeptides of the present invention can be used as targeting molecules once fused to other proteins.

Examples of domains that can be fused to polypeptides of the present invention include not only heterologous signal sequences, but also other heterologous functional regions. The fusion does not necessarily need to be direct, but may occur through linker sequences.

Moreover, fusion proteins may also be engineered to improve characteristics of the polypeptide of the present invention. For instance, a region of additional amino acids, particularly charged amino acids, may be added to the N-terminus of the polypeptide to improve stability and persistence during purification from the host cell or subsequent handling and storage. Also, peptide moieties may be added to the polypeptide to facilitate purification. Such regions may be removed prior to final preparation of the polypeptide. The addition of peptide moieties to facilitate handling of polypeptides are familiar and routine techniques in the art.

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Moreover, polypeptides of the present invention, including fragments, and specifically epitopes, can be combined with parts of the constant domain of immunoglobulins (IgG), resulting in chimeric polypeptides. These fusion proteins facilitate purification and show an increased half-life in vivo. One reported example describes chimeric proteins consisting of the first two domains of the human CD4-polypeptide and various domains of the constant regions of the heavy or light chains of mammalian immunoglobulins. (EP A 394,827; Traunecker et al., Nature 331:84-86 (1988).) Fusion proteins having disulfide-linked dimeric structures (due to the IgG) can also be more efficient in binding and neutralizing other molecules, than the monomeric secreted protein or protein fragment alone. (Fountoulakis et al., J. Biochem. 270:3958-3964 (1995).)

Similarly, EP-A-O 464 533 (Canadian counterpart 2045869) discloses fusion proteins comprising various portions of constant region of immunoglobulin molecules together with another human protein or part thereof. In many cases, the Fc part in a fusion protein is beneficial in therapy and diagnosis, and thus can result in, for example, improved pharmacokinetic properties. (EP-A 0232 262.) Alternatively, deleting the Fc part after the fusion protein has been expressed, detected, and purified, would be desired. For example, the Fc portion may hinder therapy and diagnosis if the fusion protein is used as an antigen for immunizations. In drug discovery, for example, human proteins, such as hIL-5, have been fused with Fc portions for the purpose of high-throughput screening assays to identify antagonists of hIL-5. (See, D. Bennett et al., J. Molecular Recognition 8:52-58 (1995); K. Johanson et al., J. Biol. Chem. 270:9459-9471 (1995).)

Moreover, the polypeptides of the present invention can be fused to marker sequences, such as a peptide which facilitates purification of the fused polypeptide. In preferred embodiments, the marker amino acid sequence is a hexa-histidine peptide, such as the tag provided in a pQE vector (QIAGEN, Inc., 9259 Eton Avenue, Chatsworth, CA, 91311), among others, many of which are commercially available. As described in Gentz et al., Proc. Natl. Acad. Sci. USA 86:821-824 (1989), for instance, hexa-histidine provides for convenient purification of the fusion protein. Another peptide tag useful for purification, the "HA" tag, corresponds to an epitope derived from the influenza hemagglutinin protein. (Wilson et al., Cell 37:767 (1984).)

Thus, any of these above fusions can be engineered using the polynucleotides or the polypeptides of the present invention.

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Vectors, Host Cells, and Protein Production

The present invention also relates to vectors containing the polynucleotide of the present invention, host cells, and the production of polypeptides by recombinant techniques. The vector may be, for example, a phage, plasmid, viral, or retroviral vector. Retroviral vectors may be replication competent or replication defective. In the latter case, viral propagation generally will occur only in complementing host cells.

The polynucleotides may be joined to a vector containing a selectable marker for propagation in a host. Generally, a plasmid vector is introduced in a precipitate, such as a calcium phosphate precipitate, or in a complex with a charged lipid. If the vector is a virus, it may be packaged in vitro using an appropriate packaging cell line and then transduced into host cells.

The polynucleotide insert should be operatively linked to an appropriate promoter, such as the phage lambda PL promoter, the E. coli lac, trp, phoA and tac promoters, the SV40 early and late promoters and promoters of retroviral LTRs, to name a few. Other suitable promoters will be known to the skilled artisan. The expression constructs will further contain sites for transcription initiation, termination, and, in the transcribed region, a ribosome binding site for translation. The coding portion of the transcripts expressed by the constructs will preferably include a translation initiating codon at the beginning and a termination codon (UAA, UGA or UAG) appropriately positioned at the end of the polypeptide to be translated.

As indicated, the expression vectors will preferably include at least one selectable marker. Such markers include dihydrofolate reductase, G418 or neomycin resistance for eukaryotic cell culture and tetracycline, kanamycin or ampicillin resistance genes for culturing in E. coli and other bacteria. Representative examples of appropriate hosts include, but are not limited to, bacterial cells, such as E. coli, Streptomyces and Salmonella typhimurium cells; fungal cells, such as yeast cells; insect cells such as Drosophila S2 and Spodoptera Sf9 cells; animal cells such as CHO, COS, 293, and Bowes melanoma cells; and plant cells. Appropriate culture mediums and conditions for the above-described host cells are known in the art.

Among vectors preferred for use in bacteria include pQE70, pQE60 and pQE-9, available from QIAGEN, Inc.; pBluescript vectors, Phagescript vectors, pNH8A, pNH16a, pNH18A, pNH46A, available from Stratagene Cloning Systems, Inc.; and ptrc99a, pKK223-3, pKK233-3, pDR540, pRIT5 available from Pharmacia Biotech, Inc. Among preferred eukaryotic vectors are pWLNEO, pSV2CAT, pOG44, pXT1 and pSG available from Stratagene; and pSVK3, pBPV, pMSG and pSVL available from Pharmacia. Other suitable vectors will be readily apparent to the skilled artisan.

Introduction of the construct into the host cell can be effected by calcium phosphate transfection, DEAE-dextran mediated transfection, cationic lipid-mediated transfection, electroporation, transduction, infection, or other methods. Such methods are described in many standard laboratory manuals, such as Davis et al., Basic Methods In Molecular Biology (1986). It is specifically contemplated that the polypeptides of the present invention may in fact be expressed by a host cell lacking a recombinant vector.

A polypeptide of this invention can be recovered and purified from recombinant cell cultures by well-known methods including ammonium sulfate or ethanol precipitation, acid extraction, anion or cation exchange chromatography, phosphocellulose chromatography, hydrophobic interaction chromatography, affinity chromatography, hydroxylapatite chromatography and lectin chromatography. Most preferably, high performance liquid chromatography ("HPLC") is employed for purification.

Polypeptides of the present invention, and preferably the secreted form, can also be recovered from: products purified from natural sources, including bodily fluids, tissues and cells, whether directly isolated or cultured; products of chemical synthetic procedures; and products produced by recombinant techniques from a prokaryotic or eukaryotic host, including, for example, bacterial, yeast, higher plant, insect, and mammalian cells. Depending upon the host employed in a recombinant production procedure, the polypeptides of the present invention may be glycosylated or may be non-glycosylated. In addition, polypeptides of the invention may also include an initial modified methionine residue, in some cases as a result of host-mediated processes. Thus, it is well known in the art that the N-terminal methionine encoded by the translation initiation codon generally is removed with high efficiency from any protein after translation in all eukaryotic cells. While the N-terminal methionine on most proteins also is efficiently removed in most prokaryotes, for some proteins, this prokaryotic removal process is inefficient, depending on the nature of the amino acid to which the N-terminal methionine is covalently linked.

30 Uses of the Polynucleotides

Each of the polynucleotides identified herein can be used in numerous ways as reagents. The following description should be considered exemplary and utilizes known techniques.

The polynucleotides of the present invention are useful for chromosome identification. There exists an ongoing need to identify new chromosome markers, since few chromosome marking reagents, based on actual sequence data (repeat

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polymorphisms), are presently available. Each polynucleotide of the present invention can be used as a chromosome marker.

Briefly, sequences can be mapped to chromosomes by preparing PCR primers (preferably 15-25 bp) from the sequences shown in SEQ ID NO:X. Primers can be selected using computer analysis so that primers do not span more than one predicted exon in the genomic DNA. These primers are then used for PCR screening of somatic cell hybrids containing individual human chromosomes. Only those hybrids containing the human gene corresponding to the SEQ ID NO:X will yield an amplified fragment.

Similarly, somatic hybrids provide a rapid method of PCR mapping the polynucleotides to particular chromosomes. Three or more clones can be assigned per day using a single thermal cycler. Moreover, sublocalization of the polynucleotides can be achieved with panels of specific chromosome fragments. Other gene mapping strategies that can be used include in situ hybridization, prescreening with labeled flow-sorted chromosomes, and preselection by hybridization to construct chromosome specific-cDNA libraries.

Precise chromosomal location of the polynucleotides can also be achieved using fluorescence in situ hybridization (FISH) of a metaphase chromosomal spread. This technique uses polynucleotides as short as 500 or 600 bases; however, polynucleotides 2,000-4,000 bp are preferred. For a review of this technique, see Verma et al., "Human Chromosomes: a Manual of Basic Techniques," Pergamon Press, New York (1988).

For chromosome mapping, the polynucleotides can be used individually (to mark a single chromosome or a single site on that chromosome) or in panels (for marking multiple sites and/or multiple chromosomes). Preferred polynucleotides correspond to the noncoding regions of the cDNAs because the coding sequences are more likely conserved within gene families, thus increasing the chance of cross hybridization during chromosomal mapping.

Once a polynucleotide has been mapped to a precise chromosomal location, the physical position of the polynucleotide can be used in linkage analysis. Linkage analysis establishes coinheritance between a chromosomal location and presentation of a particular disease. (Disease mapping data are found, for example, in V. McKusick, Mendelian Inheritance in Man (available on line through Johns Hopkins University Welch Medical Library).) Assuming 1 megabase mapping resolution and one gene per 20 kb, a cDNA precisely localized to a chromosomal region associated with the disease could be one of 50-500 potential causative genes.

Thus, once coinheritance is established, differences in the polynucleotide and the corresponding gene between affected and unaffected individuals can be examined.

First, visible structural alterations in the chromosomes, such as deletions or translocations, are examined in chromosome spreads or by PCR. If no structural alterations exist, the presence of point mutations are ascertained. Mutations observed in some or all affected individuals, but not in normal individuals, indicates that the mutation may cause the disease. However, complete sequencing of the polypeptide and the corresponding gene from several normal individuals is required to distinguish the mutation from a polymorphism. If a new polymorphism is identified, this polymorphic polypeptide can be used for further linkage analysis.

Furthermore, increased or decreased expression of the gene in affected individuals as compared to unaffected individuals can be assessed using polynucleotides of the present invention. Any of these alterations (altered expression, chromosomal rearrangement, or mutation) can be used as a diagnostic or prognostic marker.

In addition to the foregoing, a polynucleotide can be used to control gene expression through triple helix formation or antisense DNA or RNA. Both methods 15 rely on binding of the polynucleotide to DNA or RNA. For these techniques, preferred polynucleotides are usually 20 to 40 bases in length and complementary to either the region of the gene involved in transcription (triple helix - see Lee et al., Nucl. Acids Res. 6:3073 (1979); Cooney et al., Science 241:456 (1988); and Dervan et al., Science 251:1360 (1991)) or to the mRNA itself (antisense - Okano, J. Neurochem. 56:560 20 (1991); Oligodeoxy-nucleotides as Antisense Inhibitors of Gene Expression, CRC Press, Boca Raton, FL (1988).) Triple helix formation optimally results in a shut-off of RNA transcription from DNA, while antisense RNA hybridization blocks translation of an mRNA molecule into polypeptide. Both techniques are effective in model systems, and the information disclosed herein can be used to design antisense or triple 25 helix polynucleotides in an effort to treat disease.

Polynucleotides of the present invention are also useful in gene therapy. One goal of gene therapy is to insert a normal gene into an organism having a defective gene, in an effort to correct the genetic defect. The polynucleotides disclosed in the present invention offer a means of targeting such genetic defects in a highly accurate manner. Another goal is to insert a new gene that was not present in the host genome, thereby producing a new trait in the host cell.

The polynucleotides are also useful for identifying individuals from minute biological samples. The United States military, for example, is considering the use of restriction fragment length polymorphism (RFLP) for identification of its personnel. In this technique, an individual's genomic DNA is digested with one or more restriction enzymes, and probed on a Southern blot to yield unique bands for identifying

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personnel. This method does not suffer from the current limitations of "Dog Tags" which can be lost, switched, or stolen, making positive identification difficult. The polynucleotides of the present invention can be used as additional DNA markers for RFLP.

The polynucleotides of the present invention can also be used as an alternative to RFLP, by determining the actual base-by-base DNA sequence of selected portions of an individual's genome. These sequences can be used to prepare PCR primers for amplifying and isolating such selected DNA, which can then be sequenced. Using this technique, individuals can be identified because each individual will have a unique set of DNA sequences. Once an unique ID database is established for an individual, positive identification of that individual, living or dead, can be made from extremely small tissue samples.

Forensic biology also benefits from using DNA-based identification techniques as disclosed herein. DNA sequences taken from very small biological samples such as tissues, e.g., hair or skin, or body fluids, e.g., blood, saliva, semen, etc., can be amplified using PCR. In one prior art technique, gene sequences amplified from polymorphic loci, such as DQa class II HLA gene, are used in forensic biology to identify individuals. (Erlich, H., PCR Technology, Freeman and Co. (1992).) Once these specific polymorphic loci are amplified, they are digested with one or more restriction enzymes, yielding an identifying set of bands on a Southern blot probed with DNA corresponding to the DQa class II HLA gene. Similarly, polynucleotides of the present invention can be used as polymorphic markers for forensic purposes.

There is also a need for reagents capable of identifying the source of a particular tissue. Such need arises, for example, in forensics when presented with tissue of unknown origin. Appropriate reagents can comprise, for example, DNA probes or primers specific to particular tissue prepared from the sequences of the present invention. Panels of such reagents can identify tissue by species and/or by organ type. In a similar fashion, these reagents can be used to screen tissue cultures for contamination.

In the very least, the polynucleotides of the present invention can be used as molecular weight markers on Southern gels, as diagnostic probes for the presence of a specific mRNA in a particular cell type, as a probe to "subtract-out" known sequences in the process of discovering novel polynucleotides, for selecting and making oligomers for attachment to a "gene chip" or other support, to raise anti-DNA antibodies using DNA immunization techniques, and as an antigen to elicit an immune response.

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Uses of the Polypeptides

Each of the polypeptides identified herein can be used in numerous ways. The following description should be considered exemplary and utilizes known techniques.

A polypeptide of the present invention can be used to assay protein levels in a biological sample using antibody-based techniques. For example, protein expression in tissues can be studied with classical immunohistological methods. (Jalkanen, M., et al., J. Cell. Biol. 101:976-985 (1985): Jalkanen, M., et al., J. Cell. Biol. 105:3087-3096 (1987).) Other antibody-based methods useful for detecting protein gene expression include immunoassays, such as the enzyme linked immunosorbent assay (ELISA) and the radioimmunoassay (RIA). Suitable antibody assay labels are known in the art and include enzyme labels, such as, glucose oxidase, and radioisotopes, such as iodine (1251, 1211), carbon (14C), sulfur (35S), tritium (3H), indium (112In), and technetium (99mTc), and fluorescent labels, such as fluorescein and rhodamine, and biotin.

In addition to assaying secreted protein levels in a biological sample, proteins can also be detected in vivo by imaging. Antibody labels or markers for in vivo imaging of protein include those detectable by X-radiography, NMR or ESR. For X-radiography, suitable labels include radioisotopes such as barium or cesium, which emit detectable radiation but are not overtly harmful to the subject. Suitable markers for NMR and ESR include those with a detectable characteristic spin, such as deuterium, which may be incorporated into the antibody by labeling of nutrients for the relevant hybridoma.

A protein-specific antibody or antibody fragment which has been labeled with an appropriate detectable imaging moiety, such as a radioisotope (for example, 131I, 112In, 99mTc), a radio-opaque substance, or a material detectable by nuclear magnetic resonance, is introduced (for example, parenterally, subcutaneously, or intraperitoneally) into the mammal. It will be understood in the art that the size of the subject and the imaging system used will determine the quantity of imaging moiety needed to produce diagnostic images. In the case of a radioisotope moiety, for a human subject, the quantity of radioactivity injected will normally range from about 5 to 20 millicuries of 99mTc. The labeled antibody or antibody fragment will then preferentially accumulate at the location of cells which contain the specific protein. In vivo tumor imaging is described in S.W. Burchiel et al., "Immunopharmacokinetics of Radiolabeled Antibodies and Their Fragments." (Chapter 13 in Tumor Imaging: The Radiochemical Detection of Cancer, S.W. Burchiel and B. A. Rhodes, eds., Masson Publishing Inc. (1982).)

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Thus, the invention provides a diagnostic method of a disorder, which involves (a) assaying the expression of a polypeptide of the present invention in cells or body fluid of an individual; (b) comparing the level of gene expression with a standard gene expression level, whereby an increase or decrease in the assayed polypeptide gene expression level compared to the standard expression level is indicative of a disorder.

Moreover, polypeptides of the present invention can be used to treat disease. For example, patients can be administered a polypeptide of the present invention in an effort to replace absent or decreased levels of the polypeptide (e.g., insulin), to supplement absent or decreased levels of a different polypeptide (e.g., hemoglobin S for hemoglobin B), to inhibit the activity of a polypeptide (e.g., an oncogene), to activate the activity of a polypeptide (e.g., by binding to a receptor), to reduce the activity of a membrane bound receptor by competing with it for free ligand (e.g., soluble TNF receptors used in reducing inflammation), or to bring about a desired response (e.g., blood vessel growth).

Similarly, antibodies directed to a polypeptide of the present invention can also be used to treat disease. For example, administration of an antibody directed to a polypeptide of the present invention can bind and reduce overproduction of the polypeptide. Similarly, administration of an antibody can activate the polypeptide, such as by binding to a polypeptide bound to a membrane (receptor).

At the very least, the polypeptides of the present invention can be used as molecular weight markers on SDS-PAGE gels or on molecular sieve gel filtration columns using methods well known to those of skill in the art. Polypeptides can also be used to raise antibodies, which in turn are used to measure protein expression from a recombinant cell, as a way of assessing transformation of the host cell. Moreover, the polypeptides of the present invention can be used to test the following biological activities.

Biological Activities

The polynucleotides and polypeptides of the present invention can be used in assays to test for one or more biological activities. If these polynucleotides and polypeptides do exhibit activity in a particular assay, it is likely that these molecules may be involved in the diseases associated with the biological activity. Thus, the polynucleotides and polypeptides could be used to treat the associated disease.

35 **Immune Activity**

A polypeptide or polynucleotide of the present invention may be useful in treating deficiencies or disorders of the immune system, by activating or inhibiting the

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proliferation, differentiation, or mobilization (chemotaxis) of immune cells. Immune cells develop through a process called hematopoiesis, producing myeloid (platelets, red blood cells, neutrophils, and macrophages) and lymphoid (B and T lymphocytes) cells from pluripotent stem cells. The etiology of these immune deficiencies or disorders may be genetic, somatic, such as cancer or some autoimmune disorders, acquired (e.g., by chemotherapy or toxins), or infectious. Moreover, a polynucleotide or polypeptide of the present invention can be used as a marker or detector of a particular immune system disease or disorder.

A polynucleotide or polypeptide of the present invention may be useful in treating or detecting deficiencies or disorders of hematopoietic cells. A polypeptide or polynucleotide of the present invention could be used to increase differentiation and proliferation of hematopoietic cells, including the pluripotent stem cells, in an effort to treat those disorders associated with a decrease in certain (or many) types hematopoietic cells. Examples of immunologic deficiency syndromes include, but are not limited to: blood protein disorders (e.g. agammaglobulinemia, dysgammaglobulinemia), ataxia telangiectasia, common variable immunodeficiency, Digeorge Syndrome, HIV infection, HTLV-BLV infection, leukocyte adhesion deficiency syndrome, lymphopenia, phagocyte bactericidal dysfunction, severe combined immunodeficiency (SCIDs), Wiskott-Aldrich Disorder, anemia, thrombocytopenia, or hemoglobinuria.

Moreover, a polypeptide or polynucleotide of the present invention could also be used to modulate hemostatic (the stopping of bleeding) or thrombolytic activity (clet formation). For example, by increasing hemostatic or thrombolytic activity, a polynucleotide or polypeptide of the present invention could be used to treat blood coagulation disorders (e.g., afibrinogenemia, factor deficiencies), blood platelet disorders (e.g. thrombocytopenia), or wounds resulting from trauma, surgery, or other causes. Alternatively, a polynucleotide or polypeptide of the present invention that can decrease hemostatic or thrombolytic activity could be used to inhibit or dissolve clotting. These molecules could be important in the treatment of heart attacks (infarction), strokes, or scarring.

A polynucleotide or polypeptide of the present invention may also be useful in treating or detecting autoimmune disorders. Many autoimmune disorders result from inappropriate recognition of self as foreign material by immune cells. This inappropriate recognition results in an immune response leading to the destruction of the host tissue. Therefore, the administration of a polypeptide or polynucleotide of the present invention that inhibits an immune response, particularly the proliferation, differentiation, or chemotaxis of T-cells, may be an effective therapy in preventing autoimmune disorders.

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Examples of autoimmune disorders that can be treated or detected by the present invention include, but are not limited to: Addison's Disease, hemolytic anemia. antiphospholipid syndrome, rheumatoid arthritis, dermatitis, allergic encephalomyelitis, glomerulonephritis, Goodpasture's Syndrome, Graves' Disease, Multiple Sclerosis. Myasthenia Gravis, Neuritis, Ophthalmia, Bullous Pemphigoid, Pemphigus, Polyendocrinopathies, Purpura, Reiter's Disease, Stiff-Man Syndrome, Autoimmune Thyroiditis, Systemic Lupus Erythematosus, Autoimmune Pulmonary Inflammation, Guillain-Barre Syndrome, insulin dependent diabetes mellitis, and autoimmune inflammatory eye disease.

Similarly, allergic reactions and conditions, such as asthma (particularly allergic asthma) or other respiratory problems, may also be treated by a polypeptide or polynucleotide of the present invention. Moreover, these molecules can be used to treat anaphylaxis, hypersensitivity to an antigenic molecule, or blood group incompatibility.

A polynucleotide or polypeptide of the present invention may also be used to treat and/or prevent organ rejection or graft-versus-host disease (GVHD). Organ rejection occurs by host immune cell destruction of the transplanted tissue through an immune response. Similarly, an immune response is also involved in GVHD, but, in this case, the foreign transplanted immune cells destroy the host tissues. The administration of a polypeptide or polynucleotide of the present invention that inhibits an immune response, particularly the proliferation, differentiation, or chemotaxis of T-cells, may be an effective therapy in preventing organ rejection or GVHD.

Similarly, a polypeptide or polynucleotide of the present invention may also be used to modulate inflammation. For example, the polypeptide or polynucleotide may inhibit the proliferation and differentiation of cells involved in an inflammatory response. These molecules can be used to treat inflammatory conditions, both chronic and acute conditions, including inflammation associated with infection (e.g., septic shock, sepsis, or systemic inflammatory response syndrome (SIRS)), ischemia-reperfusion injury, endotoxin lethality, arthritis, complement-mediated hyperacute rejection, nephritis, cytokine or chemokine induced lung injury, inflammatory bowel disease, Crohn's disease, or resulting from over production of cytokines (e.g., TNF or IL-1.)

Hyperproliferative Disorders

A polypeptide or polynucleotide can be used to treat or detect hyperproliferative disorders, including neoplasms. A polypeptide or polynucleotide of the present invention may inhibit the proliferation of the disorder through direct or indirect

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interactions. Alternatively, a polypeptide or polynucleotide of the present invention may proliferate other cells which can inhibit the hyperproliferative disorder.

For example, by increasing an immune response, particularly increasing antigenic qualities of the hyperproliferative disorder or by proliferating, differentiating, or mobilizing T-cells, hyperproliferative disorders can be treated. This immune response may be increased by either enhancing an existing immune response, or by initiating a new immune response. Alternatively, decreasing an immune response may also be a method of treating hyperproliferative disorders, such as a chemotherapeutic agent.

Examples of hyperproliferative disorders that can be treated or detected by a polynucleotide or polypeptide of the present invention include, but are not limited to neoplasms located in the: abdomen, bone, breast, digestive system, liver, pancreas, peritoneum, endocrine glands (adrenal, parathyroid, pituitary, testicles, ovary, thymus, thyroid), eye, head and neck, nervous (central and peripheral), lymphatic system, pelvic, skin, soft tissue, spleen, thoracic, and urogenital.

Similarly, other hyperproliferative disorders can also be treated or detected by a polynucleotide or polypeptide of the present invention. Examples of such hyperproliferative disorders include, but are not limited to: hypergammaglobulinemia, lymphoproliferative disorders, paraproteinemias, purpura, sarcoidosis, Sezary Syndrome, Waldenstron's Macroglobulinemia, Gaucher's Disease, histiocytosis, and any other hyperproliferative disease, besides neoplasia, located in an organ system listed above.

Infectious Disease

A polypeptide or polynucleotide of the present invention can be used to treat or detect infectious agents. For example, by increasing the immune response, particularly increasing the proliferation and differentiation of B and/or T cells, infectious diseases may be treated. The immune response may be increased by either enhancing an existing immune response, or by initiating a new immune response. Alternatively, the polypeptide or polynucleotide of the present invention may also directly inhibit the infectious agent, without necessarily eliciting an immune response.

Viruses are one example of an infectious agent that can cause disease or symptoms that can be treated or detected by a polynucleotide or polypeptide of the present invention. Examples of viruses, include, but are not limited to the following DNA and RNA viral families: Arbovirus, Adenoviridae, Arenaviridae, Arterivirus, Birnaviridae, Bunyaviridae, Caliciviridae, Circoviridae, Coronaviridae, Flaviviridae, Hepadnaviridae (Hepatitis), Herpesviridae (such as, Cytomegalovirus, Herpes

Simplex. Herpes Zoster), Mononegavirus (e.g., Paramyxoviridae, Morbillivirus, Rhabdoviridae), Orthomyxoviridae (e.g., Influenza), Papovaviridae, Parvoviridae, Picomaviridae, Poxviridae (such as Smallpox or Vaccinia), Reoviridae (e.g., Rotavirus), Retroviridae (HTLV-I, HTLV-II, Lentivirus), and Togaviridae (e.g., Rubivirus). Viruses falling within these families can cause a variety of diseases or symptoms, including, but not limited to: arthritis, bronchiollitis, encephalitis, eye infections (e.g., conjunctivitis, keratitis), chronic fatigue syndrome, hepatitis (A, B, C, E, Chronic Active, Delta), meningitis, opportunistic infections (e.g., AIDS), pneumonia, Burkitt's Lymphoma, chickenpox, hemorrhagic fever, Measles, Mumps,
Parainfluenza, Rabies, the common cold, Polio, leukemia, Rubella, sexually transmitted diseases, skin diseases (e.g., Kaposi's, warts), and viremia. A polypeptide or polynucleotide of the present invention can be used to treat or detect any of these symptoms or diseases.

Similarly, bacterial or fungal agents that can cause disease or symptoms and that 15 can be treated or detected by a polynucleotide or polypeptide of the present invention include, but not limited to, the following Gram-Negative and Gram-positive bacterial families and fungi: Actinomycetales (e.g., Corynebacterium, Mycobacterium, Norcardia), Aspergillosis, Bacillaceae (e.g., Anthrax, Clostridium), Bacteroidaceae, Blastomycosis, Bordetella, Borrelia, Brucellosis, Candidiasis, Campylobacter, 20 Coccidioidomycosis, Cryptococcosis, Dermatocycoses, Enterobacteriaceae (Klebsiella, Salmonella, Serratia, Yersinia), Erysipelothrix, Helicobacter, Legionellosis, Leptospirosis, Listeria, Mycoplasmatales, Neisseriaceae (e.g., Acinetobacter, Gonorrhea, Menigococcal), Pasteurellacea Infections (e.g., Actinobacillus, Heamophilus, Pasteurella), Pseudomonas, Rickettsiaceae, Chlamydiaceae, Syphilis, 25 and Staphylococcal. These bacterial or fungal families can cause the following diseases or symptoms, including, but not limited to: bacteremia, endocarditis, eye infections (conjunctivitis, tuberculosis, uveitis), gingivitis, opportunistic infections (e.g., AIDS related infections), paronychia, prosthesis-related infections, Reiter's Disease, respiratory tract infections, such as Whooping Cough or Empyema, sepsis, Lyme 30 Disease, Cat-Scratch Disease, Dysentery, Paratyphoid Fever, food poisoning, Typhoid, pneumonia, Gonorrhea, meningitis, Chlamydia, Syphilis, Diphtheria, Leprosy, Paratuberculosis, Tuberculosis, Lupus, Botulism, gangrene, tetanus, impetigo, Rheumatic Fever, Scarlet Fever, sexually transmitted diseases, skin diseases (e.g., cellulitis, dermatocycoses), toxemia, urinary tract infections, wound infections. 35 A polypeptide or polynucleotide of the present invention can be used to treat or detect

any of these symptoms or diseases.

Moreover, parasitic agents causing disease or symptoms that can be treated or detected by a polynucleotide or polypeptide of the present invention include, but not limited to, the following families: Amebiasis, Babesiosis, Coccidiosis, Cryptosporidiosis, Dientamoebiasis, Dourine, Ectoparasitic, Giardiasis, Helminthiasis, Leishmaniasis, Theileriasis, Toxoplasmosis, Trypanosomiasis, and Trichomonas. These parasites can cause a variety of diseases or symptoms, including, but not limited to: Scabies, Trombiculiasis, eye infections, intestinal disease (e.g., dysentery, giardiasis), liver disease, lung disease, opportunistic infections (e.g., AIDS related), Malaria, pregnancy complications, and toxoplasmosis. A polypeptide or polynucleotide of the present invention can be used to treat or detect any of these symptoms or diseases.

Preferably, treatment using a polypeptide or polynucleotide of the present invention could either be by administering an effective amount of a polypeptide to the patient, or by removing cells from the patient, supplying the cells with a polynucleotide of the present invention, and returning the engineered cells to the patient (ex vivo therapy). Moreover, the polypeptide or polynucleotide of the present invention can be used as an antigen in a vaccine to raise an immune response against infectious disease.

Regeneration

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A polynucleotide or polypeptide of the present invention can be used to differentiate, proliferate, and attract cells, leading to the regeneration of tissues. (See, Science 276:59-87 (1997).) The regeneration of tissues could be used to repair, replace, or protect tissue damaged by congenital defects, trauma (wounds, burns, incisions, or ulcers), age, disease (e.g. osteoporosis, osteocarthritis, periodontal disease, liver failure), surgery, including cosmetic plastic surgery, fibrosis, reperfusion injury, or systemic cytokine damage.

Tissues that could be regenerated using the present invention include organs (e.g., pancreas, liver, intestine, kidney, skin, endothelium), muscle (smooth, skeletal or cardiac), vascular (including vascular endothelium), nervous, hematopoietic, and skeletal (bone, cartilage, tendon, and ligament) tissue. Preferably, regeneration occurs without or decreased scarring. Regeneration also may include angiogenesis.

Moreover, a polynucleotide or polypeptide of the present invention may increase regeneration of tissues difficult to heal. For example, increased tendon/ligament regeneration would quicken recovery time after damage. A polynucleotide or polypeptide of the present invention could also be used prophylactically in an effort to avoid damage. Specific diseases that could be treated include of tendinitis, carpal tunnel syndrome, and other tendon or ligament defects. A further example of tissue

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regeneration of non-healing wounds includes pressure ulcers, ulcers associated with vascular insufficiency, surgical, and traumatic wounds.

Similarly, nerve and brain tissue could also be regenerated by using a polynucleotide or polypeptide of the present invention to proliferate and differentiate nerve cells. Diseases that could be treated using this method include central and peripheral nervous system diseases, neuropathies, or mechanical and traumatic disorders (e.g., spinal cord disorders, head trauma, cerebrovascular disease, and stoke). Specifically, diseases associated with peripheral nerve injuries, peripheral neuropathy (e.g., resulting from chemotherapy or other medical therapies), localized neuropathies, and central nervous system diseases (e.g., Alzheimer's disease, Parkinson's disease, Huntington's disease, amyotrophic lateral sclerosis, and Shy-Drager syndrome), could all be treated using the polynucleotide or polypeptide of the present invention.

15 **Chemotaxis**

A polynucleotide or polypeptide of the present invention may have chemotaxis activity. A chemotaxic molecule attracts or mobilizes cells (e.g., monocytes, fibroblasts, neutrophils, T-cells, mast cells, eosinophils, epithelial and/or endothelial cells) to a particular site in the body, such as inflammation, infection, or site of hyperproliferation. The mobilized cells can then fight off and/or heal the particular trauma or abnormality.

A polynucleotide or polypeptide of the present invention may increase chemotaxic activity of particular cells. These chemotactic molecules can then be used to treat inflammation, infection, hyperproliferative disorders, or any immune system disorder by increasing the number of cells targeted to a particular location in the body. For example, chemotaxic molecules can be used to treat wounds and other trauma to tissues by attracting immune cells to the injured location. Chemotactic molecules of the present invention can also attract fibroblasts, which can be used to treat wounds.

It is also contemplated that a polynucleotide or polypeptide of the present invention may inhibit chemotactic activity. These molecules could also be used to treat disorders. Thus, a polynucleotide or polypeptide of the present invention could be used as an inhibitor of chemotaxis.

Binding Activity

A polypeptide of the present invention may be used to screen for molecules that bind to the polypeptide or for molecules to which the polypeptide binds. The binding of the polypeptide and the molecule may activate (agonist), increase, inhibit

(antagonist), or decrease activity of the polypeptide or the molecule bound. Examples of such molecules include antibodies, oligonucleotides, proteins (e.g., receptors), or small molecules.

Preferably, the molecule is closely related to the natural ligand of the polypeptide, e.g., a fragment of the ligand, or a natural substrate, a ligand, a structural or functional mimetic. (See, Coligan et al., Current Protocols in Immunology 1(2):Chapter 5 (1991).) Similarly, the molecule can be closely related to the natural receptor to which the polypeptide binds, or at least, a fragment of the receptor capable of being bound by the polypeptide (e.g., active site). In either case, the molecule can be rationally designed using known techniques.

Preferably, the screening for these molecules involves producing appropriate cells which express the polypeptide, either as a secreted protein or on the cell membrane. Preferred cells include cells from mammals, yeast, Drosophila, or *E. coli*. Cells expressing the polypeptide (or cell membrane containing the expressed polypeptide) are then preferably contacted with a test compound potentially containing the molecule to observe binding, stimulation, or inhibition of activity of either the polypeptide or the molecule.

The assay may simply test binding of a candidate compound to the polypeptide, wherein binding is detected by a label, or in an assay involving competition with a labeled competitor. Further, the assay may test whether the candidate compound results in a signal generated by binding to the polypeptide.

Alternatively, the assay can be carried out using cell-free preparations, polypeptide/molecule affixed to a solid support, chemical libraries, or natural product mixtures. The assay may also simply comprise the steps of mixing a candidate compound with a solution containing a polypeptide, measuring polypeptide/molecule activity or binding, and comparing the polypeptide/molecule activity or binding to a standard.

Preferably, an ELISA assay can measure polypeptide level or activity in a sample (e.g., biological sample) using a monoclonal or polyclonal antibody. The antibody can measure polypeptide level or activity by either binding, directly or indirectly, to the polypeptide or by competing with the polypeptide for a substrate.

All of these above assays can be used as diagnostic or prognostic markers. The molecules discovered using these assays can be used to treat disease or to bring about a particular result in a patient (e.g., blood vessel growth) by activating or inhibiting the polypeptide/molecule. Moreover, the assays can discover agents which may inhibit or enhance the production of the polypeptide from suitably manipulated cells or tissues.

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Therefore, the invention includes a method of identifying compounds which bind to a polypeptide of the invention comprising the steps of: (a) incubating a candidate binding compound with a polypeptide of the invention; and (b) determining if binding has occurred. Moreover, the invention includes a method of identifying agonists/antagonists comprising the steps of: (a) incubating a candidate compound with a polypeptide of the invention, (b) assaying a biological activity, and (b) determining if a biological activity of the polypeptide has been altered.

Other Activities

A polypeptide or polynucleotide of the present invention may also increase or decrease the differentiation or proliferation of embryonic stem cells, besides, as discussed above, hematopoietic lineage.

A polypeptide or polynucleotide of the present invention may also be used to modulate mammalian characteristics, such as body height, weight, hair color, eye color, skin. percentage of adipose tissue, pigmentation, size, and shape (e.g., cosmetic surgery). Similarly, a polypeptide or polynucleotide of the present invention may be used to modulate mammalian metabolism affecting catabolism, anabolism, processing, utilization, and storage of energy.

A polypeptide or polynucleotide of the present invention may be used to change a mammal's mental state or physical state by influencing biorhythms, caricadic rhythms, depression (including depressive disorders), tendency for violence, tolerance for pain, reproductive capabilities (preferably by Activin or Inhibin-like activity), hormonal or endocrine levels, appetite, libido, memory, stress, or other cognitive qualities.

A polypeptide or polynucleotide of the present invention may also be used as a food additive or preservative, such as to increase or decrease storage capabilities, fat content, lipid, protein, carbohydrate, vitamins, minerals, cofactors or other nutritional components.

30 Other Preferred Embodiments

Other preferred embodiments of the claimed invention include an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to a sequence of at least about 50 contiguous nucleotides in the nucleotide sequence of SEQ ID NO:X wherein X is any integer as defined in Table 1.

Also preferred is a nucleic acid molecule wherein said sequence of contiguous nucleotides is included in the nucleotide sequence of SEQ ID NO:X in the range of

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positions beginning with the nucleotide at about the position of the 5' Nucleotide of the Clone Sequence and ending with the nucleotide at about the position of the 3' Nucleotide of the Clone Sequence as defined for SEQ ID NO:X in Table 1.

Also preferred is a nucleic acid molecule wherein said sequence of contiguous nucleotides is included in the nucleotide sequence of SEQ ID NO:X in the range of positions beginning with the nucleotide at about the position of the 5' Nucleotide of the Start Codon and ending with the nucleotide at about the position of the 3' Nucleotide of the Clone Sequence as defined for SEQ ID NO:X in Table 1.

Similarly preferred is a nucleic acid molecule wherein said sequence of contiguous nucleotides is included in the nucleotide sequence of SEQ ID NO:X in the range of positions beginning with the nucleotide at about the position of the 5' Nucleotide of the First Amino Acid of the Signal Peptide and ending with the nucleotide at about the position of the 3' Nucleotide of the Clone Sequence as defined for SEQ ID NO:X in Table 1.

Also preferred is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to a sequence of at least about 150 contiguous nucleotides in the nucleotide sequence of SEQ ID NO:X.

Further preferred is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to a sequence of at least about 500 contiguous nucleotides in the nucleotide sequence of SEQ ID NO:X.

A further preferred embodiment is a nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to the nucleotide sequence of SEQ ID NO:X beginning with the nucleotide at about the position of the 5' Nucleotide of the First Amino Acid of the Signal Peptide and ending with the nucleotide at about the position of the 3' Nucleotide of the Clone Sequence as defined for SEQ ID NO:X in Table 1.

A further preferred embodiment is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to the complete nucleotide sequence of SEQ ID NO:X.

Also preferred is an isolated nucleic acid molecule which hybridizes under stringent hybridization conditions to a nucleic acid molecule, wherein said nucleic acid molecule which hybridizes does not hybridize under stringent hybridization conditions to a nucleic acid molecule having a nucleotide sequence consisting of only A residues or of only T residues.

Also preferred is a composition of matter comprising a DNA molecule which comprises a human cDNA clone identified by a cDNA Clone Identifier in Table 1, which DNA molecule is contained in the material deposited with the American Type

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Culture Collection and given the ATCC Deposit Number shown in Table 1 for said cDNA Clone Identifier.

Also preferred is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to a sequence of at least 50 contiguous nucleotides in the nucleotide sequence of a human cDNA clone identified by a cDNA Clone Identifier in Table 1, which DNA molecule is contained in the deposit given the ATCC Deposit Number shown in Table 1.

Also preferred is an isolated nucleic acid molecule, wherein said sequence of at least 50 contiguous nucleotides is included in the nucleotide sequence of the complete open reading frame sequence encoded by said human cDNA clone.

Also preferred is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to sequence of at least 150 contiguous nucleotides in the nucleotide sequence encoded by said human cDNA clone.

A further preferred embodiment is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to sequence of at least 500 contiguous nucleotides in the nucleotide sequence encoded by said human cDNA clone.

A further preferred embodiment is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to the complete nucleotide sequence encoded by said human cDNA clone.

A further preferred embodiment is a method for detecting in a biological sample a nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to a sequence of at least 50 contiguous nucleotides in a sequence selected from the group consisting of: a nucleotide sequence of SEQ ID NO:X wherein X is any integer as defined in Table 1; and a nucleotide sequence encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with-the ATCC Deposit Number shown for said cDNA clone in Table 1; which method comprises a step of comparing a nucleotide sequence of at least one nucleic acid molecule in said sample with a sequence selected from said group and determining whether the sequence of said nucleic acid molecule in said sample is at least 95% identical to said selected sequence.

Also preferred is the above method wherein said step of comparing sequences comprises determining the extent of nucleic acid hybridization between nucleic acid molecules in said sample and a nucleic acid molecule comprising said sequence selected from said group. Similarly, also preferred is the above method wherein said step of comparing sequences is performed by comparing the nucleotide sequence determined from a nucleic acid molecule in said sample with said sequence selected from said group. The nucleic acid molecules can comprise DNA molecules or RNA molecules.

A further preferred embodiment is a method for identifying the species, tissue or cell type of a biological sample which method comprises a step of detecting nucleic acid molecules in said sample, if any, comprising a nucleotide sequence that is at least 95% identical to a sequence of at least 50 contiguous nucleotides in a sequence selected from the group consisting of: a nucleotide sequence of SEQ ID NO:X wherein X is any integer as defined in Table 1; and a nucleotide sequence encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

The method for identifying the species, tissue or cell type of a biological sample can comprise a step of detecting nucleic acid molecules comprising a nucleotide sequence in a panel of at least two nucleotide sequences, wherein at least one sequence in said panel is at least 95% identical to a sequence of at least 50 contiguous nucleotides in a sequence selected from said group.

Also preferred is a method for diagnosing in a subject a pathological condition associated with abnormal structure or expression of a gene encoding a secreted protein identified in Table 1, which method comprises a step of detecting in a biological sample obtained from said subject nucleic acid molecules, if any, comprising a nucleotide sequence that is at least 95% identical to a sequence of at least 50 contiguous nucleotides in a sequence selected from the group consisting of: a nucleotide sequence of SEQ ID NO:X wherein X is any integer as defined in Table 1; and a nucleotide sequence encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

The method for diagnosing a pathological condition can comprise a step of detecting nucleic acid molecules comprising a nucleotide sequence in a panel of at least two nucleotide sequences, wherein at least one sequence in said panel is at least 95% identical to a sequence of at least 50 contiguous nucleotides in a sequence selected from said group.

Also preferred is a composition of matter comprising isolated nucleic acid molecules wherein the nucleotide sequences of said nucleic acid molecules comprise a panel of at least two nucleotide sequences, wherein at least one sequence in said panel is at least 95% identical to a sequence of at least 50 contiguous nucleotides in a sequence selected from the group consisting of: a nucleotide sequence of SEQ ID NO:X wherein X is any integer as defined in Table 1; and a nucleotide sequence encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1. The nucleic acid molecules can comprise DNA molecules or RNA molecules.

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Also preferred is an isolated polypeptide comprising an amino acid sequence at least 90% identical to a sequence of at least about 10 contiguous amino acids in the amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1.

Also preferred is a polypeptide, wherein said sequence of contiguous amino acids is included in the amino acid sequence of SEQ ID NO:Y in the range of positions beginning with the residue at about the position of the First Amino Acid of the Secreted Portion and ending with the residue at about the Last Amino Acid of the Open Reading Frame as set forth for SEQ ID NO:Y in Table 1.

Also preferred is an isolated polypeptide comprising an amino acid sequence at least 95% identical to a sequence of at least about 30 contiguous amino acids in the amino acid sequence of SEQ ID NO:Y.

Further preferred is an isolated polypeptide comprising an amino acid sequence at least 95% identical to a sequence of at least about 100 contiguous amino acids in the amino acid sequence of SEQ ID NO:Y.

Further preferred is an isolated polypeptide comprising an amino acid sequence at least 95% identical to the complete amino acid sequence of SEQ ID NO:Y.

Further preferred is an isolated polypeptide comprising an amino acid sequence at least 90% identical to a sequence of at least about 10 contiguous amino acids in the complete amino acid sequence of a secreted protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Also preferred is a polypeptide wherein said sequence of contiguous amino acids is included in the amino acid sequence of a secreted portion of the secreted protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Also preferred is an isolated polypeptide comprising an amino acid sequence at least 95% identical to a sequence of at least about 30 contiguous amino acids in the amino acid sequence of the secreted portion of the protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Also preferred is an isolated polypeptide comprising an amino acid sequence at least 95% identical to a sequence of at least about 100 contiguous amino acids in the amino acid sequence of the secreted portion of the protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

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Also preferred is an isolated polypeptide comprising an amino acid sequence at least 95% identical to the amino acid sequence of the secreted portion of the protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Further preferred is an isolated antibody which binds specifically to a polypeptide comprising an amino acid sequence that is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Further preferred is a method for detecting in a biological sample a polypeptide comprising an amino acid sequence which is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1; which method comprises a step of comparing an amino acid sequence of at least one polypeptide molecule in said sample with a sequence selected from said group and determining whether the sequence of said polypeptide molecule in said sample is at least 90% identical to said sequence of at least 10 contiguous amino acids.

Also preferred is the above method wherein said step of comparing an amino acid sequence of at least one polypeptide molecule in said sample with a sequence selected from said group comprises determining the extent of specific binding of polypeptides in said sample to an antibody which binds specifically to a polypeptide comprising an amino acid sequence that is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Also preferred is the above method wherein said step of comparing sequences is performed by comparing the amino acid sequence determined from a polypeptide molecule in said sample with said sequence selected from said group.

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Also preferred is a method for identifying the species, tissue or cell type of a biological sample which method comprises a step of detecting polypeptide molecules in said sample, if any, comprising an amino acid sequence that is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a secreted protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Also preferred is the above method for identifying the species, tissue or cell type of a biological sample, which method comprises a step of detecting polypeptide molecules comprising an amino acid sequence in a panel of at least two amino acid sequences, wherein at least one sequence in said panel is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the above group.

Also preferred is a method for diagnosing in a subject a pathological condition associated with abnormal structure or expression of a gene encoding a secreted protein identified in Table 1, which method comprises a step of detecting in a biological sample obtained from said subject polypeptide molecules comprising an amino acid sequence in a panel of at least two amino acid sequences, wherein at least one sequence in said panel is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a secreted protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

In any of these methods, the step of detecting said polypeptide molecules includes using an antibody.

Also preferred is an isolated nucleic acid molecule comprising a nucleotide sequence which is at least 95% identical to a nucleotide sequence encoding a polypeptide wherein said polypeptide comprises an amino acid sequence that is at least 90% identical to a sequence of at least 10 contiguous amino acids in a sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a secreted protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

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Also preferred is an isolated nucleic acid molecule, wherein said nucleotide sequence encoding a polypeptide has been optimized for expression of said polypeptide in a prokaryotic host.

Also preferred is an isolated nucleic acid molecule, wherein said polypeptide comprises an amino acid sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y wherein Y is any integer as defined in Table 1; and a complete amino acid sequence of a secreted protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1.

Further preferred is a method of making a recombinant vector comprising inserting any of the above isolated nucleic acid molecule into a vector. Also preferred is the recombinant vector produced by this method. Also preferred is a method of making a recombinant host cell comprising introducing the vector into a host cell, as well as the recombinant host cell produced by this method.

Also preferred is a method of making an isolated polypeptide comprising culturing this recombinant host cell under conditions such that said polypeptide is expressed and recovering said polypeptide. Also preferred is this method of making an isolated polypeptide, wherein said recombinant host cell is a eukaryotic cell and said polypeptide is a secreted portion of a human secreted protein comprising an amino acid sequence selected from the group consisting of: an amino acid sequence of SEQ ID NO:Y beginning with the residue at the position of the First Amino Acid of the Secreted Portion of SEQ ID NO:Y wherein Y is an integer set forth in Table 1 and said position of the First Amino Acid of the Secreted Portion of SEQ ID NO:Y is defined in Table 1; and an amino acid sequence of a secreted portion of a protein encoded by a human cDNA clone identified by a cDNA Clone Identifier in Table 1 and contained in the deposit with the ATCC Deposit Number shown for said cDNA clone in Table 1. The isolated polypeptide produced by this method is also preferred.

Also preferred is a method of treatment of an individual in need of an increased level of a secreted protein activity, which method comprises administering to such an individual a pharmaceutical composition comprising an amount of an isolated polypeptide, polynucleotide, or antibody of the claimed invention effective to increase the level of said protein activity in said individual.

Having generally described the invention, the same will be more readily understood by reference to the following examples, which are provided by way of illustration and are not intended as limiting.

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Examples

Example 1: Isolation of a Selected cDNA Clone From the Deposited Sample

Each cDNA clone in a cited ATCC deposit is contained in a plasmid vector. Table 1 identifies the vectors used to construct the cDNA library from which each clone was isolated. In many cases, the vector used to construct the library is a phage vector from which a plasmid has been excised. The table immediately below correlates the related plasmid for each phage vector used in constructing the cDNA library. For example, where a particular clone is identified in Table 1 as being isolated in the vector "Lambda Zap," the corresponding deposited clone is in "pBluescript."

	Vector Used to Construct Library	Corresponding Deposited Plasmid
	Lambda Zap	pBluescript (pBS)
	Uni-Zap XR	pBluescript (pBS)
15	Zap Express	pBK
	lafmid BA	plafmid BA
	pSport1	pSport I
	pCMVSport 2.0	pCMVSport 2.0
	pCMVSport 3.0	pCMVSport 3.0
20	pCR [®] 2.1	pCR [®] 2.1

Vectors Lambda Zap (U.S. Patent Nos. 5,128,256 and 5,286,636), Uni-Zap XR (U.S. Patent Nos. 5,128, 256 and 5,286,636), Zap Express (U.S. Patent Nos. 5,128,256 and 5,286,636), pBluescript (pBS) (Short, J. M. et al., Nucleic Acids Res. 16:7583-7600 (1988); Alting-Mees, M. A. and Short, J. M., Nucleic Acids Res. 17:9494 (1989)) and pBK (Alting-Mees, M. A. et al., Strategies 5:58-61 (1992)) are

- commercially available from Stratagene Cloning Systems, Inc., 11011 N. Torrey Pines Road, La Jolla, CA, 92037. pBS contains an ampicillin resistance gene and pBK contains a neomycin resistance gene. Both can be transformed into E. coli strain XL-1 Blue, also available from Stratagene. pBS comes in 4 forms SK+, SK-, KS+ and KS.
- The S and K refers to the orientation of the polylinker to the T7 and T3 primer sequences which flank the polylinker region ("S" is for SacI and "K" is for KpnI which are the first sites on each respective end of the linker). "+" or "-" refer to the orientation of the f1 origin of replication ("ori"), such that in one orientation, single stranded rescue initiated from the f1 ori generates sense strand DNA and in the other, antisense.

Vectors pSport1, pCMVSport 2.0 and pCMVSport 3.0, were obtained from Life Technologies, Inc., P. O. Box 6009, Gaithersburg, MD 20897. All Sport vectors contain an ampicillin resistance gene and may be transformed into E. coli strain

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DH10B, also available from Life Technologies. (See, for instance, Gruber, C. E., et al., Focus 15:59 (1993).) Vector lafmid BA (Bento Soares, Columbia University, NY) contains an ampicillin resistance gene and can be transformed into E. coli strain XL-1 Blue. Vector pCR®2.1, which is available from Invitrogen, 1600 Faraday Avenue, Carlsbad, CA 92008, contains an ampicillin resistance gene and may be transformed into E. coli strain DH10B, available from Life Technologies. (See, for instance, Clark, J. M., Nuc. Acids Res. 16:9677-9686 (1988) and Mead, D. et al., Bio/Technology 9: (1991).) Preferably, a polynucleotide of the present invention does not comprise the phage vector sequences identified for the particular clone in Table 1, as well as the corresponding plasmid vector sequences designated above.

The deposited material in the sample assigned the ATCC Deposit Number cited in Table 1 for any given cDNA clone also may contain one or more additional plasmids, each comprising a cDNA clone different from that given clone. Thus, deposits sharing the same ATCC Deposit Number contain at least a plasmid for each cDNA clone identified in Table 1. Typically, each ATCC deposit sample cited in Table 1 comprises a mixture of approximately equal amounts (by weight) of about 50 plasmid DNAs, each containing a different cDNA clone; but such a deposit sample may include plasmids for more or less than 50 cDNA clones, up to about 500 cDNA clones.

Two approaches can be used to isolate a particular clone from the deposited sample of plasmid DNAs cited for that clone in Table 1. First, a plasmid is directly isolated by screening the clones using a polynucleotide probe corresponding to SEQ ID NO:X.

Particularly, a specific polynucleotide with 30-40 nucleotides is synthesized

using an Applied Biosystems DNA synthesizer according to the sequence reported.

The oligonucleotide is labeled, for instance, with ³²P-γ-ATP using T4 polynucleotide kinase and purified according to routine methods. (E.g., Maniatis et al., Molecular Cloning: A Laboratory Manual, Cold Spring Harbor Press, Cold Spring, NY (1982).)

The plasmid mixture is transformed into a suitable host, as indicated above (such as XL-1 Blue (Stratagene)) using techniques known to those of skill in the art, such as those provided by the vector supplier or in related publications or patents cited above. The transformants are plated on 1.5% agar plates (containing the appropriate selection agent, e.g., ampicillin) to a density of about 150 transformants (colonies) per plate. These plates are screened using Nylon membranes according to routine methods for

Manual, 2nd Edit., (1989), Cold Spring Harbor Laboratory Press, pages 1.93 to 1.104), or other techniques known to those of skill in the art.

bacterial colony screening (e.g., Sambrook et al., Molecular Cloning: A Laboratory

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Alternatively, two primers of 17-20 nucleotides derived from both ends of the SEQ ID NO:X (i.e., within the region of SEQ ID NO:X bounded by the 5' NT and the 3' NT of the clone defined in Table 1) are synthesized and used to amplify the desired cDNA using the deposited cDNA plasmid as a template. The polymerase chain reaction is carried out under routine conditions, for instance, in 25 µl of reaction mixture with 0.5 ug of the above cDNA template. A convenient reaction mixture is 1.5-5 mM MgCl₂, 0.01% (w/v) gelatin, 20 µM each of dATP, dCTP, dGTP, dTTP, 25 pmol of each primer and 0.25 Unit of Taq polymerase. Thirty five cycles of PCR (denaturation at 94°C for 1 min; annealing at 55°C for 1 min; elongation at 72°C for 1 min) are performed with a Perkin-Elmer Cetus automated thermal cycler. The amplified product is analyzed by agarose gel electrophoresis and the DNA band with expected molecular weight is excised and purified. The PCR product is verified to be the selected sequence by subcloning and sequencing the DNA product.

Several methods are available for the identification of the 5' or 3' non-coding portions of a gene which may not be present in the deposited clone. These methods include but are not limited to, filter probing, clone enrichment using specific probes, and protocols similar or identical to 5' and 3' "RACE" protocols which are well known in the art. For instance, a method similar to 5' RACE is available for generating the missing 5' end of a desired full-length transcript. (Fromont-Racine et al., Nucleic Acids Res. 21(7):1683-1684 (1993).)

Briefly, a specific RNA oligonucleotide is ligated to the 5' ends of a population of RNA presumably containing full-length gene RNA transcripts. A primer set containing a primer specific to the ligated RNA oligonucleotide and a primer specific to a known sequence of the gene of interest is used to PCR amplify the 5' portion of the desired full-length gene. This amplified product may then be sequenced and used to generate the full length gene.

This above method starts with total RNA isolated from the desired source, although poly-A+ RNA can be used. The RNA preparation can then be treated with phosphatase if necessary to eliminate 5' phosphate groups on degraded or damaged RNA which may interfere with the later RNA ligase step. The phosphatase should then be inactivated and the RNA treated with tobacco acid pyrophosphatase in order to remove the cap structure present at the 5' ends of messenger RNAs. This reaction leaves a 5' phosphate group at the 5' end of the cap cleaved RNA which can then be ligated to an RNA oligonucleotide using T4 RNA ligase.

This modified RNA preparation is used as a template for first strand cDNA synthesis using a gene specific oligonucleotide. The first strand synthesis reaction is

used as a template for PCR amplification of the desired 5' end using a primer specific to the ligated RNA oligonucleotide and a primer specific to the known sequence of the gene of interest. The resultant product is then sequenced and analyzed to confirm that the 5' end sequence belongs to the desired gene.

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Example 2: Isolation of Genomic Clones Corresponding to a Polynucleotide

A human genomic P1 library (Genomic Systems, Inc.) is screened by PCR using primers selected for the cDNA sequence corresponding to SEQ ID NO:X., according to the method described in Example 1. (See also, Sambrook.)

Example 3: Tissue Distribution of Polypeptide

Tissue distribution of mRNA expression of polynucleotides of the present invention is determined using protocols for Northern blot analysis, described by, among others, Sambrook et al. For example, a cDNA probe produced by the method described in Example 1 is labeled with P³² using the rediprimeTM DNA labeling system (Amersham Life Science), according to manufacturer's instructions. After labeling, the probe is purified using CHROMA SPIN-100TM column (Clontech Laboratories, Inc.), according to manufacturer's protocol number PT1200-1. The purified labeled probe is then used to examine various human tissues for mRNA expression.

Multiple Tissue Northern (MTN) blots containing various human tissues (H) or human immune system tissues (IM) (Clontech) are examined with the labeled probe using ExpressHybTM hybridization solution (Clontech) according to manufacturer's protocol number PT1190-1. Following hybridization and washing, the blots are mounted and exposed to film at -70°C overnight, and the films developed according to standard procedures.

Example 4: Chromosomal Mapping of the Polynucleotides

An oligonucleotide primer set is designed according to the sequence at the 5' end of SEQ ID NO:X. This primer preferably spans about 100 nucleotides. This primer set is then used in a polymerase chain reaction under the following set of conditions: 30 seconds, 95°C; 1 minute, 56°C; 1 minute, 70°C. This cycle is repeated 32 times followed by one 5 minute cycle at 70°C. Human, mouse, and hamster DNA is used as template in addition to a somatic cell hybrid panel containing individual chromosomes or chromosome fragments (Bios, Inc). The reactions is analyzed on

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either 8% polyacrylamide gels or 3.5 % agarose gels. Chromosome mapping is determined by the presence of an approximately 100 bp PCR fragment in the particular somatic cell hybrid.

5 Example 5: Bacterial Expression of a Polypeptide

A polynucleotide encoding a polypeptide of the present invention is amplified using PCR oligonucleotide primers corresponding to the 5' and 3' ends of the DNA sequence, as outlined in Example 1, to synthesize insertion fragments. The primers used to amplify the cDNA insert should preferably contain restriction sites, such as BamHI and XbaI, at the 5' end of the primers in order to clone the amplified product into the expression vector. For example, BamHI and XbaI correspond to the restriction enzyme sites on the bacterial expression vector pQE-9. (Qiagen, Inc., Chatsworth, CA). This plasmid vector encodes antibiotic resistance (Amp^T), a bacterial origin of replication (ori), an IPTG-regulatable promoter/operator (P/O), a ribosome binding site (RBS), a 6-histidine tag (6-His), and restriction enzyme cloning sites.

The pQE-9 vector is digested with BamHI and XbaI and the amplified fragment is ligated into the pQE-9 vector maintaining the reading frame initiated at the bacterial RBS. The ligation mixture is then used to transform the E. coli strain M15/rep4 (Qiagen, Inc.) which contains multiple copies of the plasmid pREP4, which expresses the lacI repressor and also confers kanamycin resistance (Kan^r). Transformants are identified by their ability to grow on LB plates and ampicillin/kanamycin resistant colonies are selected. Plasmid DNA is isolated and confirmed by restriction analysis.

Clones containing the desired constructs are grown overnight (O/N) in liquid culture in LB media supplemented with both Amp (100 ug/ml) and Kan (25 ug/ml). The O/N culture is used to inoculate a large culture at a ratio of 1:100 to 1:250. The cells are grown to an optical density 600 (O.D.⁶⁰⁰) of between 0.4 and 0.6. IPTG (Isopropyl-B-D-thiogalacto pyranoside) is then added to a final concentration of 1 mM. IPTG induces by inactivating the lacI repressor, clearing the P/O leading to increased gene expression.

Cells are grown for an extra 3 to 4 hours. Cells are then harvested by centrifugation (20 mins at 6000Xg). The cell pellet is solubilized in the chaotropic agent 6 Molar Guanidine HCl by stirring for 3-4 hours at 4°C. The cell debris is removed by centrifugation, and the supernatant containing the polypeptide is loaded onto a nickel-nitrilo-tri-acetic acid ("Ni-NTA") affinity resin column (available from QIAGEN, Inc., *supra*). Proteins with a 6 x His tag bind to the Ni-NTA resin with high

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affinity and can be purified in a simple one-step procedure (for details see: The QIAexpressionist (1995) QIAGEN, Inc., *supra*).

Briefly, the supernatant is loaded onto the column in 6 M guanidine-HCl, pH 8, the column is first washed with 10 volumes of 6 M guanidine-HCl, pH 8, then washed with 10 volumes of 6 M guanidine-HCl pH 6, and finally the polypeptide is eluted with 6 M guanidine-HCl, pH 5.

The purified protein is then renatured by dialyzing it against phosphate-buffered saline (PBS) or 50 mM Na-acetate, pH 6 buffer plus 200 mM NaCl. Alternatively, the protein can be successfully refolded while immobilized on the Ni-NTA column. The recommended conditions are as follows: renature using a linear 6M-1M urea gradient in 500 mM NaCl, 20% glycerol, 20 mM Tris/HCl pH 7.4, containing protease inhibitors. The renaturation should be performed over a period of 1.5 hours or more. After renaturation the proteins are eluted by the addition of 250 mM immidazole. Immidazole is removed by a final dialyzing step against PBS or 50 mM sodium acetate pH 6 buffer plus 200 mM NaCl. The purified protein is stored at 4°C or frozen at -80°C.

In addition to the above expression vector, the present invention further includes an expression vector comprising phage operator and promoter elements operatively linked to a polynucleotide of the present invention, called pHE4a. (ATCC Accession Number 209645, deposited on February 25, 1998.) This vector contains: 1) a neomycinphosphotransferase gene as a selection marker, 2) an E. coli origin of replication, 3) a T5 phage promoter sequence, 4) two lac operator sequences, 5) a Shine-Delgarno sequence, and 6) the lactose operon repressor gene (lacIq). The origin of replication (oriC) is derived from pUC19 (LTI, Gaithersburg, MD). The promoter sequence and operator sequences are made synthetically.

DNA can be inserted into the pHEa by restricting the vector with NdeI and XbaI, BamHI, XhoI, or Asp718, running the restricted product on a gel, and isolating the larger fragment (the stuffer fragment should be about 310 base pairs). The DNA insert is generated according to the PCR protocol described in Example 1, using PCR primers having restriction sites for NdeI (5' primer) and XbaI, BamHI, XhoI, or Asp718 (3' primer). The PCR insert is gel purified and restricted with compatible enzymes. The insert and vector are ligated according to standard protocols.

The engineered vector could easily be substituted in the above protocol to express protein in a bacterial system.

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Example 6: Purification of a Polypeptide from an Inclusion Body

The following alternative method can be used to purify a polypeptide expressed in $E \, coli$ when it is present in the form of inclusion bodies. Unless otherwise specified, all of the following steps are conducted at 4-10°C.

Upon completion of the production phase of the *E. coli* fermentation, the cell culture is cooled to 4-10°C and the cells harvested by continuous centrifugation at 15,000 rpm (Heraeus Sepatech). On the basis of the expected yield of protein per unit weight of cell paste and the amount of purified protein required, an appropriate amount of cell paste, by weight, is suspended in a buffer solution containing 100 mM Tris, 50 mM EDTA, pH 7.4. The cells are dispersed to a homogeneous suspension using a high shear mixer.

The cells are then lysed by passing the solution through a microfluidizer (Microfuidics, Corp. or APV Gaulin, Inc.) twice at 4000-6000 psi. The homogenate is then mixed with NaCl solution to a final concentration of 0.5 M NaCl, followed by centrifugation at 7000 xg for 15 min. The resultant pellet is washed again using 0.5M NaCl, 100 mM Tris, 50 mM EDTA, pH 7.4.

The resulting washed inclusion bodies are solubilized with 1.5 M guanidine hydrochloride (GuHCl) for 2-4 hours. After 7000 xg centrifugation for 15 min., the pellet is discarded and the polypeptide containing supernatant is incubated at 4°C overnight to allow further GuHCl extraction.

Following high speed centrifugation (30,000 xg) to remove insoluble particles, the GuHCl solubilized protein is refolded by quickly mixing the GuHCl extract with 20 volumes of buffer containing 50 mM sodium, pH 4.5, 150 mM NaCl, 2 mM EDTA by vigorous stirring. The refolded diluted protein solution is kept at 4°C without mixing for 12 hours prior to further purification steps.

To clarify the refolded polypeptide solution, a previously prepared tangential filtration unit equipped with 0.16 µm membrane filter with appropriate surface area (e.g., Filtron), equilibrated with 40 mM sodium acetate, pH 6.0 is employed. The filtered sample is loaded onto a cation exchange resin (e.g., Poros HS-50, Perseptive Biosystems). The column is washed with 40 mM sodium acetate, pH 6.0 and eluted with 250 mM, 500 mM, 1000 mM, and 1500 mM NaCl in the same buffer, in a stepwise manner. The absorbance at 280 nm of the effluent is continuously monitored. Fractions are collected and further analyzed by SDS-PAGE.

Fractions containing the polypeptide are then pooled and mixed with 4 volumes of water. The diluted sample is then loaded onto a previously prepared set of tandem columns of strong anion (Poros HQ-50, Perseptive Biosystems) and weak anion (Poros CM-20, Perseptive Biosystems) exchange resins. The columns are equilibrated with 40 mM sodium acetate, pH 6.0. Both columns are washed with 40 mM sodium acetate, pH 6.0, 200 mM NaCl. The CM-20 column is then eluted using a 10 column volume linear gradient ranging from 0.2 M NaCl, 50 mM sodium acetate, pH 6.0 to 1.0 M NaCl, 50 mM sodium acetate, pH 6.5. Fractions are collected under constant A₂₈₀ monitoring of the effluent. Fractions containing the polypeptide (determined, for instance, by 16% SDS-PAGE) are then pooled.

The resultant polypeptide should exhibit greater than 95% purity after the above refolding and purification steps. No major contaminant bands should be observed from Commassie blue stained 16% SDS-PAGE gel when 5 μ g of purified protein is loaded. The purified protein can also be tested for endotoxin/LPS contamination, and typically the LPS content is less than 0.1 ng/ml according to LAL assays.

Example 7: Cloning and Expression of a Polypeptide in a Baculovirus Expression System

In this example, the plasmid shuttle vector pA2 is used to insert a polynucleotide into a baculovirus to express a polypeptide. This expression vector contains the strong polyhedrin promoter of the *Autographa californica* nuclear polyhedrosis virus (AcMNPV) followed by convenient restriction sites such as BamHI, Xba I and Asp718. The polyadenylation site of the simian virus 40 ("SV40") is used for efficient polyadenylation. For easy selection of recombinant virus, the plasmid contains the beta-galactosidase gene from *E. coli* under control of a weak Drosophila promoter in the same orientation, followed by the polyadenylation signal of the polyhedrin gene. The inserted genes are flanked on both sides by viral sequences for cell-mediated homologous recombination with wild-type viral DNA to generate a viable virus that express the cloned polynucleotide.

Many other baculovirus vectors can be used in place of the vector above, such as pAc373, pVL941, and pAcIM1, as one skilled in the art would readily appreciate, as long as the construct provides appropriately located signals for transcription, translation, secretion and the like, including a signal peptide and an in-frame AUG as required. Such vectors are described, for instance, in Luckow et al., Virology 170:31-39 (1989).

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Specifically, the cDNA sequence contained in the deposited clone, including the AUG initiation codon and the naturally associated leader sequence identified in Table 1, is amplified using the PCR protocol described in Example 1. If the naturally occurring signal sequence is used to produce the secreted protein, the pA2 vector does not need a second signal peptide. Alternatively, the vector can be modified (pA2 GP) to include a baculovirus leader sequence, using the standard methods described in Summers et al., "A Manual of Methods for Baculovirus Vectors and Insect Cell Culture Procedures," Texas Agricultural Experimental Station Bulletin No. 1555 (1987).

The amplified fragment is isolated from a 1% agarose gel using a commercially available kit ("Geneclean," BIO 101 Inc., La Jolla, Ca.). The fragment then is digested with appropriate restriction enzymes and again purified on a 1% agarose gel.

The plasmid is digested with the corresponding restriction enzymes and optionally, can be dephosphorylated using calf intestinal phosphatase, using routine procedures known in the art. The DNA is then isolated from a 1% agarose gel using a commercially available kit ("Geneclean" BIO 101 Inc., La Jolla, Ca.).

The fragment and the dephosphorylated plasmid are ligated together with T4 DNA ligase. *E. coli* HB101 or other suitable *E. coli* hosts such as XL-1 Blue (Stratagene Cloning Systems, La Jolla, CA) cells are transformed with the ligation mixture and spread on culture plates. Bacteria containing the plasmid are identified by digesting DNA from individual colonies and analyzing the digestion product by gel electrophoresis. The sequence of the cloned fragment is confirmed by DNA sequencing.

Five μg of a plasmid containing the polynucleotide is co-transfected with 1.0 μg of a commercially available linearized baculovirus DNA ("BaculoGoldTM baculovirus DNA", Pharmingen, San Diego, CA), using the lipofection method described by Felgner et al., Proc. Natl. Acad. Sci. USA 84:7413-7417 (1987). One μg of BaculoGoldTM virus DNA and 5 μg of the plasmid are mixed in a sterile well of a microtiter plate containing 50 μl of serum-free Grace's medium (Life Technologies Inc., Gaithersburg, MD). Afterwards, 10 μl Lipofectin plus 90 μl Grace's medium are added, mixed and incubated for 15 minutes at room temperature. Then the transfection mixture is added drop-wise to Sf9 insect cells (ATCC CRL 1711) seeded in a 35 mm tissue culture plate with 1 ml Grace's medium without serum. The plate is then incubated for 5 hours at 27° C. The transfection solution is then removed from the plate and 1 ml of Grace's insect medium supplemented with 10% fetal calf serum is added. Cultivation is then continued at 27° C for four days.

After four days the supernatant is collected and a plaque assay is performed, as described by Summers and Smith, *supra*. An agarose gel with "Blue Gal" (Life

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Technologies Inc., Gaithersburg) is used to allow easy identification and isolation of gal-expressing clones, which produce blue-stained plaques. (A detailed description of a "plaque assay" of this type can also be found in the user's guide for insect cell culture and baculovirology distributed by Life Technologies Inc., Gaithersburg, page 9-10.) After appropriate incubation, blue stained plaques are picked with the tip of a micropipettor (e.g., Eppendorf). The agar containing the recombinant viruses is then resuspended in a microcentrifuge tube containing 200 µl of Grace's medium and the suspension containing the recombinant baculovirus is used to infect Sf9 cells seeded in 35 mm dishes. Four days later the supernatants of these culture dishes are harvested and then they are stored at 4° C.

To verify the expression of the polypeptide, Sf9 cells are grown in Grace's medium supplemented with 10% heat-inactivated FBS. The cells are infected with the recombinant baculovirus containing the polynucleotide at a multiplicity of infection ("MOI") of about 2. If radiolabeled proteins are desired, 6 hours later the medium is removed and is replaced with SF900 II medium minus methionine and cysteine (available from Life Technologies Inc., Rockville, MD). After 42 hours, 5 μ Ci of ³⁵S-methionine and 5 μ Ci ³⁵S-cysteine</sup> (available from Amersham) are added. The cells are further incubated for 16 hours and then are harvested by centrifugation. The proteins in the supernatant as well as the intracellular proteins are analyzed by SDS-PAGE followed by autoradiography (if radiolabeled).

Microsequencing of the amino acid sequence of the amino terminus of purified protein may be used to determine the amino terminal sequence of the produced protein.

Example 8: Expression of a Polypeptide in Mammalian Cells

The polypeptide of the present invention can be expressed in a mammalian cell.

A typical mammalian expression vector contains a promoter element, which mediates the initiation of transcription of mRNA, a protein coding sequence, and signals required for the termination of transcription and polyadenylation of the transcript. Additional elements include enhancers, Kozak sequences and intervening sequences flanked by donor and acceptor sites for RNA splicing. Highly efficient transcription is achieved with the early and late promoters from SV40, the long terminal repeats (LTRs) from Retroviruses, e.g., RSV, HTLVI, HIVI and the early promoter of the cytomegalovirus (CMV). However, cellular elements can also be used (e.g., the human actin promoter).

Suitable expression vectors for use in practicing the present invention include, for example, vectors such as pSVL and pMSG (Pharmacia, Uppsala, Sweden), pRSVcat (ATCC 37152), pSV2dhfr (ATCC 37146), pBC12MI (ATCC 67109), pCMVSport 2.0, and pCMVSport 3.0. Mammalian host cells that could be used

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include, human Hela, 293, H9 and Jurkat cells, mouse NIH3T3 and C127 cells, Cos 1, Cos 7 and CV1, quail QC1-3 cells, mouse L cells and Chinese hamster ovary (CHO) cells.

Alternatively, the polypeptide can be expressed in stable cell lines containing the polynucleotide integrated into a chromosome. The co-transfection with a selectable marker such as dhfr, gpt, neomycin, hygromycin allows the identification and isolation of the transfected cells.

The transfected gene can also be amplified to express large amounts of the encoded protein. The DHFR (dihydrofolate reductase) marker is useful in developing cell lines that carry several hundred or even several thousand copies of the gene of interest. (See, e.g., Alt, F. W., et al., J. Biol. Chem. 253:1357-1370 (1978); Hamlin, J. L. and Ma, C., Biochem. et Biophys. Acta, 1097:107-143 (1990); Page, M. J. and Sydenham, M. A., Biotechnology 9:64-68 (1991).) Another useful selection marker is the enzyme glutamine synthase (GS) (Murphy et al., Biochem J. 227:277-279 (1991); Bebbington et al., Bio/Technology 10:169-175 (1992). Using these markers, the mammalian cells are grown in selective medium and the cells with the highest resistance are selected. These cell lines contain the amplified gene(s) integrated into a chromosome. Chinese hamster ovary (CHO) and NSO cells are often used for the production of proteins.

Derivatives of the plasmid pSV2-dhfr (ATCC Accession No. 37146), the expression vectors pC4 (ATCC Accession No. 209646) and pC6 (ATCC Accession No.209647) contain the strong promoter (LTR) of the Rous Sarcoma Virus (Cullen et al., Molecular and Cellular Biology, 438-447 (March, 1985)) plus a fragment of the CMV-enhancer (Boshart et al., Cell 41:521-530 (1985).) Multiple cloning sites, e.g., with the restriction enzyme cleavage sites BamHI, XbaI and Asp718, facilitate the cloning of the gene of interest. The vectors also contain the 3' intron, the polyadenylation and termination signal of the rat preproinsulin gene, and the mouse DHFR gene under control of the SV40 early promoter.

Specifically, the plasmid pC6, for example, is digested with appropriate restriction enzymes and then dephosphorylated using calf intestinal phosphates by procedures known in the art. The vector is then isolated from a 1% agarose gel.

A polynucleotide of the present invention is amplified according to the protocol outlined in Example 1. If the naturally occurring signal sequence is used to produce the secreted protein, the vector does not need a second signal peptide. Alternatively, if the naturally occurring signal sequence is not used, the vector can be modified to include a heterologous signal sequence. (See, e.g., WO 96/34891.)

The amplified fragment is isolated from a 1% agarose gel using a commercially available kit ("Geneclean," BIO 101 Inc., La Jolla, Ca.). The fragment then is digested with appropriate restriction enzymes and again purified on a 1% agarose gel.

The amplified fragment is then digested with the same restriction enzyme and purified on a 1% agarose gel. The isolated fragment and the dephosphorylated vector are then ligated with T4 DNA ligase. *E. coli* HB101 or XL-1 Blue cells are then transformed and bacteria are identified that contain the fragment inserted into plasmid pC6 using, for instance, restriction enzyme analysis.

Chinese hamster ovary cells lacking an active DHFR gene is used for transfection. Five µg of the expression plasmid pC6 is cotransfected with 0.5 µg of the plasmid pSVneo using lipofectin (Felgner et al., supra). The plasmid pSV2-neo contains a dominant selectable marker, the neo gene from Tn5 encoding an enzyme that confers resistance to a group of antibiotics including G418. The cells are seeded in alpha minus MEM supplemented with 1 mg/ml G418. After 2 days, the cells are trypsinized and seeded in hybridoma cloning plates (Greiner, Germany) in alpha minus MEM supplemented with 10, 25, or 50 ng/ml of metothrexate plus 1 mg/ml G418. After about 10-14 days single clones are trypsinized and then seeded in 6-well petri dishes or 10 ml flasks using different concentrations of methotrexate (50 nM, 100 nM, 200 nM, 400 nM, 800 nM). Clones growing at the highest concentrations of methotrexate are then transferred to new 6-well plates containing even higher concentrations of methotrexate (1 μM , 2 μM , 5 μM , 10 mM, 20 mM). The same procedure is repeated until clones are obtained which grow at a concentration of 100 -200 µM. Expression of the desired gene product is analyzed, for instance, by SDS-PAGE and Western blot or by reversed phase HPLC analysis.

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Example 9: Protein Fusions

These fusion proteins can be used for a variety of applications. For example, fusion of the present polypeptides to His-tag, HA-tag, protein A, IgG domains, and maltose binding protein facilitates purification. (See Example 5; see also EP A 394,827; Traunecker, et al., Nature 331:84-86 (1988).) Similarly, fusion to IgG-1, IgG-3, and albumin increases the halflife time in vivo. Nuclear localization signals fused to the polypeptides of the present invention can target the protein to a specific subcellular localization, while covalent heterodimer or homodimers can increase or decrease the activity of a fusion protein. Fusion proteins can also create chimeric molecules having more than one function. Finally, fusion proteins can increase solubility and/or stability of the fused protein compared to the non-fused protein. All of the types of fusion

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proteins described above can be made by modifying the following protocol, which outlines the fusion of a polypeptide to an IgG molecule, or the protocol described in Example 5.

Briefly, the human Fc portion of the IgG molecule can be PCR amplified, using primers that span the 5' and 3' ends of the sequence described below. These primers also should have convenient restriction enzyme sites that will facilitate cloning into an expression vector, preferably a mammalian expression vector.

For example, if pC4 (Accession No. 209646) is used, the human Fc portion can be ligated into the BamHI cloning site. Note that the 3' BamHI site should be destroyed. Next, the vector containing the human Fc portion is re-restricted with BamHI, linearizing the vector, and a polynucleotide of the present invention, isolated by the PCR protocol described in Example 1, is ligated into this BamHI site. Note that the polynucleotide is cloned without a stop codon, otherwise a fusion protein will not be produced.

If the naturally occurring signal sequence is used to produce the secreted protein, pC4 does not need a second signal peptide. Alternatively, if the naturally occurring signal sequence is not used, the vector can be modified to include a heterologous signal sequence. (See, e.g., WO 96/34891.)

20 Human IgG Fc region:

GGGATCCGGAGCCCAAATCTTCTGACAAAACTCACACATGCCCACCGTGCC CAGCACCTGAATTCGAGGGTGCACCGTCAGTCTTCCTCTTCCCCCAAAACC CAAGGACACCCTCATGATCTCCCGGACTCCTGAGGTCACATGCGTGGTGGT GGACGTAAGCCACGAAGACCCTGAGGTCAAGTTCAACTGGTACGTGGACG 25 GCGTGGAGGTGCATAATGCCAAGACAAAGCCGCGGGAGGAGCAGTACAAC AGCACGTACCGTGTGGTCAGCGTCCTCACCGTCCTGCACCAGGACTGGCTG AATGGCAAGGAGTACAAGTGCAAGGTCTCCAACAAGCCCTCCCAACCCCC ATCGAGAAAACCATCTCCAAAGCCAAAGGGCAGCCCCGAGAACCACAGGT GTACACCCTGCCCCATCCCGGGATGAGCTGACCAAGAACCAGGTCAGCCT 30 GACCTGCCTGGTCAAAGGCTTCTATCCAAGCGACATCGCCGTGGAGTGGGA GAGCAATGGCCAGCCGGAGAACAACTACAAGACCACGCCTCCCGTGCTGG ACTCCGACGCTCCTTCTTCCTCTACAGCAAGCTCACCGTGGACAAGAGCA GGTGGCAGCAGGGAACGTCTTCTCATGCTCCGTGATGCATGAGGCTCTGC ACAACCACTACACGCAGAAGAGCCTCTCCCTGTCTCCGGGTAAATGAGTGC GACGGCCGCGACTCTAGAGGAT (SEQ ID NO:1) 35

Example 10: Production of an Antibody from a Polypeptide

The antibodies of the present invention can be prepared by a variety of methods. (See, Current Protocols, Chapter 2.) For example, cells expressing a polypeptide of the present invention is administered to an animal to induce the production of sera containing polyclonal antibodies. In a preferred method, a preparation of the secreted protein is prepared and purified to render it substantially free of natural contaminants. Such a preparation is then introduced into an animal in order to produce polyclonal antisera of greater specific activity.

In the most preferred method, the antibodies of the present invention are monoclonal antibodies (or protein binding fragments thereof). Such monoclonal antibodies can be prepared using hybridoma technology. (Köhler et al., Nature 256:495 (1975); Köhler et al., Eur. J. Immunol. 6:511 (1976); Köhler et al., Eur. J. Immunol. 6:292 (1976); Hammerling et al., in: Monoclonal Antibodies and T-Cell Hybridomas, Elsevier, N.Y., pp. 563-681 (1981).) In general, such procedures involve immunizing an animal (preferably a mouse) with polypeptide or, more preferably, with a secreted polypeptide-expressing cell. Such cells may be cultured in any suitable tissue culture medium; however, it is preferable to culture cells in Earle's modified Eagle's medium supplemented with 10% fetal bovine serum (inactivated at about 56°C), and supplemented with about 10 g/l of nonessential amino acids, about 1,000 U/ml of penicillin, and about 100 μg/ml of streptomycin.

The splenocytes of such mice are extracted and fused with a suitable myeloma cell line. Any suitable myeloma cell line may be employed in accordance with the present invention; however, it is preferable to employ the parent myeloma cell line (SP2O), available from the ATCC. After fusion, the resulting hybridoma cells are selectively maintained in HAT medium, and then cloned by limiting dilution as described by Wands et al. (Gastroenterology 80:225-232 (1981).) The hybridoma cells obtained through such a selection are then assayed to identify clones which secrete antibodies capable of binding the polypeptide.

Alternatively, additional antibodies capable of binding to the polypeptide can be produced in a two-step procedure using anti-idiotypic antibodies. Such a method makes use of the fact that antibodies are themselves antigens, and therefore, it is possible to obtain an antibody which binds to a second antibody. In accordance with this method, protein specific antibodies are used to immunize an animal, preferably a mouse. The splenocytes of such an animal are then used to produce hybridoma cells, and the hybridoma cells are screened to identify clones which produce an antibody whose ability to bind to the protein-specific antibody can be blocked by the polypeptide.

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Such antibodies comprise anti-idiotypic antibodies to the protein-specific antibody and can be used to immunize an animal to induce formation of further protein-specific antibodies.

It will be appreciated that Fab and F(ab')2 and other fragments of the antibodies of the present invention may be used according to the methods disclosed herein. Such fragments are typically produced by proteolytic cleavage, using enzymes such as papain (to produce Fab fragments) or pepsin (to produce F(ab')2 fragments). Alternatively, secreted protein-binding fragments can be produced through the application of recombinant DNA technology or through synthetic chemistry.

For in vivo use of antibodies in humans, it may be preferable to use "humanized" chimeric monoclonal antibodies. Such antibodies can be produced using genetic constructs derived from hybridoma cells producing the monoclonal antibodies described above. Methods for producing chimeric antibodies are known in the art. (See, for review, Morrison, Science 229:1202 (1985); Oi et al., BioTechniques 4:214 (1986); Cabilly et al., U.S. Patent No. 4,816,567; Taniguchi et al., EP 171496; Morrison et al., EP 173494; Neuberger et al., WO 8601533; Robinson et al., WO 8702671; Boulianne et al., Nature 312:643 (1984); Neuberger et al., Nature 314:268 (1985).)

20 Example 11: Production Of Secreted Protein For High-Throughput Screening Assays

The following protocol produces a supernatant containing a polypeptide to be tested. This supernatant can then be used in the Screening Assays described in Examples 13-20.

First, dilute Poly-D-Lysine (644 587 Boehringer-Mannheim) stock solution (1mg/ml in PBS) 1:20 in PBS (w/o calcium or magnesium 17-516F Biowhittaker) for a working solution of 50ug/ml. Add 200 ul of this solution to each well (24 well plates) and incubate at RT for 20 minutes. Be sure to distribute the solution over each well (note: a 12-channel pipetter may be used with tips on every other channel). Aspirate off the Poly-D-Lysine solution and rinse with 1ml PBS (Phosphate Buffered Saline). The PBS should remain in the well until just prior to plating the cells and plates may be poly-lysine coated in advance for up to two weeks.

Plate 293T cells (do not carry cells past P+20) at 2 x 10⁵ cells/well in .5ml DMEM(Dulbecco's Modified Eagle Medium)(with 4.5 G/L glucose and L-glutamine (12-604F Biowhittaker))/10% heat inactivated FBS(14-503F Biowhittaker)/1x Penstrep(17-602E Biowhittaker). Let the cells grow overnight.

The next day, mix together in a sterile solution basin: 300 ul Lipofectamine (18324-012 Gibco/BRL) and 5ml Optimem I (31985070 Gibco/BRL)/96-well plate. With a small volume multi-channel pipetter, aliquot approximately 2ug of an expression vector containing a polynucleotide insert, produced by the methods described in Examples 8 or 9, into an appropriately labeled 96-well round bottom plate. With a multi-channel pipetter, add 50ul of the Lipofectamine/Optimem I mixture to each well. Pipette up and down gently to mix. Incubate at RT 15-45 minutes. After about 20 minutes, use a multi-channel pipetter to add 150ul Optimem I to each well. As a control, one plate of vector DNA lacking an insert should be transfected with each set of transfections.

Preferably, the transfection should be performed by tag-teaming the following tasks. By tag-teaming, hands on time is cut in half, and the cells do not spend too much time on PBS. First, person A aspirates off the media from four 24-well plates of cells, and then person B rinses each well with .5-1ml PBS. Person A then aspirates off PBS rinse, and person B, using a12-channel pipetter with tips on every other channel, adds the 200ul of DNA/Lipofectamine/Optimem I complex to the odd wells first, then to the even wells, to each row on the 24-well plates. Incubate at 37°C for 6 hours.

While cells are incubating, prepare appropriate media, either 1%BSA in DMEM with 1x penstrep, or CHO-5 media (116.6 mg/L of CaCl2 (anhyd); 0.00130 mg/L CuSO₄-5H₂O; 0.050 mg/L of Fe(NO₂)₃-9H₂O; 0.417 mg/L of FeSO₄-7H₂O; 311.80 20 mg/L of Kcl; 28.64 mg/L of MgCl₂; 48.84 mg/L of MgSO₄; 6995.50 mg/L of NaCl; 2400.0 mg/L of NaHCO₃; 62.50 mg/L of NaH₂PO₄-H₂0; 71.02 mg/L of Na₂HPO4; .4320 mg/L of ZnSO₄-7H₂O; .002 mg/L of Arachidonic Acid; 1.022 mg/L of Cholesterol; .070 mg/L of DL-alpha-Tocopherol-Acetate; 0.0520 mg/L of Linoleic Acid; 0.010 mg/L of Linolenic Acid; 0.010 mg/L of Myristic Acid; 0.010 mg/L of Oleic 25 Acid; 0.010 mg/L of Palmitric Acid; 0.010 mg/L of Palmitic Acid; 100 mg/L of Pluronic F-68; 0.010 mg/L of Stearic Acid; 2.20 mg/L of Tween 80; 4551 mg/L of D-Glucose; 130.85 mg/ml of L- Alanine; 147.50 mg/ml of L-Arginine-HCL; 7.50 mg/ml of L-Asparagine-H₂0; 6.65 mg/ml of L-Aspartic Acid; 29.56 mg/ml of L-Cystine-2HCL-H₂0; 31.29 mg/ml of L-Cystine-2HCL; 7.35 mg/ml of L-Glutamic Acid; 365.0 30 mg/ml of L-Glutamine; 18.75 mg/ml of Glycine; 52.48 mg/ml of L-Histidine-HCL-H₂0; 106.97 mg/ml of L-Isoleucine; 111.45 mg/ml of L-Leucine; 163.75 mg/ml of L-Lysine HCL; 32.34 mg/ml of L-Methionine; 68.48 mg/ml of L-Phenylalainine; 40.0 mg/ml of L-Proline; 26.25 mg/ml of L-Serine; 101.05 mg/ml of L-Threonine; 19.22 mg/ml of L-Tryptophan; 91.79 mg/ml of L-Tryrosine-2Na-2H₂0; 99.65 mg/ml of L-35 Valine; 0.0035 mg/L of Biotin; 3.24 mg/L of D-Ca Pantothenate; 11.78 mg/L of

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Choline Chloride; 4.65 mg/L of Folic Acid; 15.60 mg/L of i-Inositol; 3.02 mg/L of Niacinamide; 3.00 mg/L of Pyridoxal HCL; 0.031 mg/L of Pyridoxine HCL; 0.319 mg/L of Riboflavin; 3.17 mg/L of Thiamine HCL; 0.365 mg/L of Thymidine; and 0.680 mg/L of Vitamin B₁₂; 25 mM of HEPES Buffer; 2.39 mg/L of Na Hypoxanthine; 0.105 mg/L of Lipoic Acid; 0.081 mg/L of Sodium Putrescine-2HCL; 55.0 mg/L of Sodium Pyruvate; 0.0067 mg/L of Sodium Selenite; 20uM of Ethanolamine; 0.122 mg/L of Ferric Citrate; 41.70 mg/L of Methyl-B-Cyclodextrin complexed with Linoleic Acid; 33.33 mg/L of Methyl-B-Cyclodextrin complexed with Oleic Acid; and 10 mg/L of Methyl-B-Cyclodextrin complexed with Retinal) with 2mm glutamine and 1x penstrep. (BSA (81-068-3 Bayer) 100gm dissolved in 1L DMEM for a 10% BSA stock solution). Filter the media and collect 50 ul for endotoxin assay in 15ml polystyrene conical.

The transfection reaction is terminated, preferably by tag-teaming, at the end of the incubation period. Person A aspirates off the transfection media, while person B adds 1.5ml appropriate media to each well. Incubate at 37°C for 45 or 72 hours depending on the media used: 1%BSA for 45 hours or CHO-5 for 72 hours.

On day four, using a 300ul multichannel pipetter, aliquot 600ul in one 1ml deep well plate and the remaining supernatant into a 2ml deep well. The supernatants from each well can then be used in the assays described in Examples 13-20.

It is specifically understood that when activity is obtained in any of the assays described below using a supernatant, the activity originates from either the polypeptide directly (e.g., as a secreted protein) or by the polypeptide inducing expression of other proteins, which are then secreted into the supernatant. Thus, the invention further provides a method of identifying the protein in the supernatant characterized by an activity in a particular assay.

Example 12: Construction of GAS Reporter Construct

One signal transduction pathway involved in the differentiation and proliferation of cells is called the Jaks-STATs pathway. Activated proteins in the Jaks-STATs pathway bind to gamma activation site "GAS" elements or interferon-sensitive responsive element ("ISRE"), located in the promoter of many genes. The binding of a protein to these elements alter the expression of the associated gene.

GAS and ISRE elements are recognized by a class of transcription factors called Signal Transducers and Activators of Transcription, or "STATs." There are six members of the STATs family. Stat1 and Stat3 are present in many cell types, as is Stat2 (as response to IFN-alpha is widespread). Stat4 is more restricted and is not in

many cell types though it has been found in T helper class I, cells after treatment with IL-12. Stat5 was originally called mammary growth factor, but has been found at higher concentrations in other cells including myeloid cells. It can be activated in tissue culture cells by many cytokines.

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The STATs are activated to translocate from the cytoplasm to the nucleus upon tyrosine phosphorylation by a set of kinases known as the Janus Kinase ("Jaks") family. Jaks represent a distinct family of soluble tyrosine kinases and include Tyk2, Jak1, Jak2, and Jak3. These kinases display significant sequence similarity and are generally catalytically inactive in resting cells.

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The Jaks are activated by a wide range of receptors summarized in the Table below. (Adapted from review by Schidler and Darnell, Ann. Rev. Biochem. 64:621-51 (1995).) A cytokine receptor family, capable of activating Jaks, is divided into two groups: (a) Class 1 includes receptors for IL-2, IL-3, IL-4, IL-6, IL-7, IL-9, IL-11, IL-12, IL-15, Epo, PRL, GH, G-CSF, GM-CSF, LIF, CNTF, and thrombopoietin; and (b) Class 2 includes IFN-a, IFN-g, and IL-10. The Class 1 receptors share a conserved cysteine motif (a set of four conserved cysteines and one tryptophan) and a WSXWS motif (a membrane proxial region encoding Trp-Ser-Xxx-Trp-Ser (SEQ ID NO:2)).

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Thus, on binding of a ligand to a receptor, Jaks are activated, which in turn activate STATs, which then translocate and bind to GAS elements. This entire process is encompassed in the Jaks-STATs signal transduction pathway.

Therefore, activation of the Jaks-STATs pathway, reflected by the binding of the GAS or the ISRE element, can be used to indicate proteins involved in the proliferation and differentiation of cells. For example, growth factors and cytokines are known to activate the Jaks-STATs pathway. (See Table below.) Thus, by using GAS elements linked to reporter molecules, activators of the Jaks-STATs pathway can be identified.

	<u>Ligand</u>	tyk2	JAKs Jak 1	Jak2	Jak3	<u>STATS</u>	GAS(elements) or ISRE
5	IFN family IFN-a/B IFN-g II-10	+	+ + ?	- + ?	-	1,2,3 1 1,3	ISRE GAS (IRF1>Lys6>IFP)
10	gp130 family IL-6 (Pleiotrohic) Il-11(Pleiotrohic) OnM(Pleiotrohic)	+ ? ?	+ + +	+ ? +	?????	1,3 1,3 1,3	GAS (IRF1>Lys6>IFP)
15	LIF(Pleiotrohic) CNTF(Pleiotrohic) G-CSF(Pleiotrohic) IL-12(Pleiotrohic)	? -/+ ? +	+ + + -	+ + ? +	? ? ? +	1,3 1,3 1,3 1,3	
20	g-C family IL-2 (lymphocytes) IL-4 (lymph/myeloid) IL-7 (lymphocytes) IL-9 (lymphocytes) IL-13 (lymphocyte) IL-15	- - - - ?	+ + + + +	- - - ? ?	+ + + + ? +	1,3,5 6 5 5 6 5	GAS GAS (IRF1 = IFP >>Ly6)(IgH) GAS GAS GAS GAS GAS GAS
25	gp140 family IL-3 (myeloid) IL-5 (myeloid) GM-CSF (myeloid)	-	-	+ + +	- -	5 5 5	GAS (IRF1>IFP>>Ly6) GAS GAS
30 35	Growth hormone fami GH PRL EPO	ily ? ? ?	- +/- -	+ + +	- -	5 1,3,5 5	GAS(B-CAS>IRF1=IFP>>Ly6)
40	Receptor Tyrosine Kir EGF PDGF CSF-1	nases ? ? ?	+ + +	+++++	- -	1,3 1,3 1,3	GAS (IRF1) GAS (not IRF1)

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To construct a synthetic GAS containing promoter element, which is used in the Biological Assays described in Examples 13-14, a PCR based strategy is employed to generate a GAS-SV40 promoter sequence. The 5' primer contains four tandem copies of the GAS binding site found in the IRF1 promoter and previously demonstrated to bind STATs upon induction with a range of cytokines (Rothman et al., Immunity 1:457-468 (1994).), although other GAS or ISRE elements can be used instead. The 5' primer also contains 18bp of sequence complementary to the SV40 early promoter sequence and is flanked with an XhoI site. The sequence of the 5' primer is: 5':GCGCCTCGAGATTTCCCCGAAATCTAGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCGAAATGATTTCCCCCGAAATGATTTCCCCCGAAATGATTTCCCCCGAAATGATTTCCCCCGAAATGATTTCCCCCGAAATGATTTCCCCCGAAATGATTTCCCCCGAAATGATTTCCCCCGAAATATCTTGCCCATCTCAATTAG:3' (SEQ ID NO:3)

The downstream primer is complementary to the SV40 promoter and is flanked with a Hind III site: 5':GCGGCAAGCTTTTTGCAAAGCCTAGGC:3' (SEQ ID NO:4)

PCR amplification is performed using the SV40 promoter template present in the B-gal:promoter plasmid obtained from Clontech. The resulting PCR fragment is digested with XhoI/Hind III and subcloned into BLSK2-. (Stratagene.) Sequencing with forward and reverse primers confirms that the insert contains the following sequence:

With this GAS promoter element linked to the SV40 promoter, a GAS:SEAP2 reporter construct is next engineered. Here, the reporter molecule is a secreted alkaline phosphatase, or "SEAP." Clearly, however, any reporter molecule can be instead of SEAP, in this or in any of the other Examples. Well known reporter molecules that can be used instead of SEAP include chloramphenicol acetyltransferase (CAT), luciferase, alkaline phosphatase, B-galactosidase, green fluorescent protein (GFP), or any protein detectable by an antibody.

The above sequence confirmed synthetic GAS-SV40 promoter element is subcloned into the pSEAP-Promoter vector obtained from Clontech using HindIII and XhoI, effectively replacing the SV40 promoter with the amplified GAS:SV40 promoter element, to create the GAS-SEAP vector. However, this vector does not contain a neomycin resistance gene, and therefore, is not preferred for mammalian expression systems.

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Thus, in order to generate mammalian stable cell lines expressing the GAS-SEAP reporter, the GAS-SEAP cassette is removed from the GAS-SEAP vector using SalI and NotI, and inserted into a backbone vector containing the neomycin resistance gene, such as pGFP-1 (Clontech), using these restriction sites in the multiple cloning site, to create the GAS-SEAP/Neo vector. Once this vector is transfected into mammalian cells, this vector can then be used as a reporter molecule for GAS binding as described in Examples 13-14.

Other constructs can be made using the above description and replacing GAS with a different promoter sequence. For example, construction of reporter molecules containing NFK-B and EGR promoter sequences are described in Examples 15 and 16. However, many other promoters can be substituted using the protocols described in these Examples. For instance, SRE, IL-2, NFAT, or Osteocalcin promoters can be substituted, alone or in combination (e.g., GAS/NF-KB/EGR, GAS/NF-KB, Il-2/NFAT, or NF-KB/GAS). Similarly, other cell lines can be used to test reporter construct activity, such as HELA (epithelial), HUVEC (endothelial), Reh (B-cell), Saos-2 (osteoblast), HUVAC (aortic), or Cardiomyocyte.

Example 13: High-Throughput Screening Assay for T-cell Activity.

The following protocol is used to assess T-cell activity by identifying factors, such as growth factors and cytokines, that may proliferate or differentiate T-cells. T-cell activity is assessed using the GAS/SEAP/Neo construct produced in Example 12. Thus, factors that increase SEAP activity indicate the ability to activate the Jaks-STATS signal transduction pathway. The T-cell used in this assay is Jurkat T-cells (ATCC Accession No. TIB-152), although Molt-3 cells (ATCC Accession No. CRL-1552) and Molt-4 cells (ATCC Accession No. CRL-1582) cells can also be used.

Jurkat T-cells are lymphoblastic CD4+ Th1 helper cells. In order to generate stable cell lines, approximately 2 million Jurkat cells are transfected with the GAS-SEAP/neo vector using DMRIE-C (Life Technologies)(transfection procedure described below). The transfected cells are seeded to a density of approximately 20,000 cells per well and transfectants resistant to 1 mg/ml genticin selected. Resistant colonies are expanded and then tested for their response to increasing concentrations of interferon gamma. The dose response of a selected clone is demonstrated.

Specifically, the following protocol will yield sufficient cells for 75 wells containing 200 ul of cells. Thus, it is either scaled up, or performed in multiple to generate sufficient cells for multiple 96 well plates. Jurkat cells are maintained in RPMI + 10% serum with 1%Pen-Strep. Combine 2.5 mls of OPTI-MEM (Life Technologies)

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with 10 ug of plasmid DNA in a T25 flask. Add 2.5 ml OPTI-MEM containing 50 ul of DMRIE-C and incubate at room temperature for 15-45 mins.

During the incubation period, count cell concentration, spin down the required number of cells (10⁷ per transfection), and resuspend in OPTI-MEM to a final concentration of 10⁷ cells/ml. Then add 1ml of 1 x 10⁷ cells in OPTI-MEM to T25 flask and incubate at 37°C for 6 hrs. After the incubation, add 10 ml of RPMI + 15% serum.

The Jurkat:GAS-SEAP stable reporter lines are maintained in RPMI + 10% serum, 1 mg/ml Genticin, and 1% Pen-Strep. These cells are treated with supernatants containing a polypeptide as produced by the protocol described in Example 11.

On the day of treatment with the supernatant, the cells should be washed and resuspended in fresh RPMI + 10% serum to a density of 500,000 cells per ml. The exact number of cells required will depend on the number of supernatants being screened. For one 96 well plate, approximately 10 million cells (for 10 plates, 100 million cells) are required.

Transfer the cells to a triangular reservoir boat, in order to dispense the cells into a 96 well dish, using a 12 channel pipette. Using a 12 channel pipette, transfer 200 ul of cells into each well (therefore adding 100, 000 cells per well).

After all the plates have been seeded, 50 ul of the supernatants are transferred directly from the 96 well plate containing the supernatants into each well using a 12 channel pipette. In addition, a dose of exogenous interferon gamma (0.1, 1.0, 10 ng) is added to wells H9, H10, and H11 to serve as additional positive controls for the assay.

The 96 well dishes containing Jurkat cells treated with supernatants are placed in an incubator for 48 hrs (note: this time is variable between 48-72 hrs). 35 ul samples from each well are then transferred to an opaque 96 well plate using a 12 channel pipette. The opaque plates should be covered (using sellophene covers) and stored at -20°C until SEAP assays are performed according to Example 17. The plates containing the remaining treated cells are placed at 4°C and serve as a source of material for repeating the assay on a specific well if desired.

As a positive control, 100 Unit/ml interferon gamma can be used which is known to activate Jurkat T cells. Over 30 fold induction is typically observed in the positive control wells.

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Example 14: High-Throughput Screening Assay Identifying Myeloid Activity

The following protocol is used to assess myeloid activity by identifying factors, such as growth factors and cytokines, that may proliferate or differentiate myeloid cells. Myeloid cell activity is assessed using the GAS/SEAP/Neo construct produced in Example 12. Thus, factors that increase SEAP activity indicate the ability to activate the Jaks-STATS signal transduction pathway. The myeloid cell used in this assay is U937, a pre-monocyte cell line, although TF-1, HL60, or KG1 can be used.

To transiently transfect U937 cells with the GAS/SEAP/Neo construct produced in Example 12, a DEAE-Dextran method (Kharbanda et. al., 1994, Cell Growth & Differentiation, 5:259-265) is used. First, harvest 2x10e⁷ U937 cells and wash with PBS. The U937 cells are usually grown in RPMI 1640 medium containing 10% heat-inactivated fetal bovine serum (FBS) supplemented with 100 units/ml penicillin and 100 mg/ml streptomycin.

Next, suspend the cells in 1 ml of 20 mM Tris-HCl (pH 7.4) buffer containing 0.5 mg/ml DEAE-Dextran, 8 ug GAS-SEAP2 plasmid DNA, 140 mM NaCl, 5 mM KCl, 375 uM Na₂HPO₄.7H₂O, 1 mM MgCl₂, and 675 uM CaCl₂. Incubate at 37°C for 45 min.

Wash the cells with RPMI 1640 medium containing 10% FBS and then resuspend in 10 ml complete medium and incubate at 37°C for 36 hr.

The GAS-SEAP/U937 stable cells are obtained by growing the cells in 400 ug/ml G418. The G418-free medium is used for routine growth but every one to two months, the cells should be re-grown in 400 ug/ml G418 for couple of passages.

These cells are tested by harvesting $1x10^8$ cells (this is enough for ten 96-well plates assay) and wash with PBS. Suspend the cells in 200 ml above described growth medium, with a final density of $5x10^5$ cells/ml. Plate 200 ul cells per well in the 96-well plate (or $1x10^5$ cells/well).

Add 50 ul of the supernatant prepared by the protocol described in Example 11. Incubate at 37°C for 48 to 72 hr. As a positive control, 100 Unit/ml interferon gamma can be used which is known to activate U937 cells. Over 30 fold induction is typically observed in the positive control wells. SEAP assay the supernatant according to the protocol described in Example 17.

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Example 15: High-Throughput Screening Assay Identifying Neuronal Activity.

When cells undergo differentiation and proliferation, a group of genes are activated through many different signal transduction pathways. One of these genes, EGR1 (early growth response gene 1), is induced in various tissues and cell types upon activation. The promoter of EGR1 is responsible for such induction. Using the EGR1 promoter linked to reporter molecules, activation of cells can be assessed.

Particularly, the following protocol is used to assess neuronal activity in PC12 cell lines. PC12 cells (rat phenochromocytoma cells) are known to proliferate and/or differentiate by activation with a number of mitogens, such as TPA (tetradecanoyl phorbol acetate), NGF (nerve growth factor), and EGF (epidermal growth factor). The EGR1 gene expression is activated during this treatment. Thus, by stably transfecting PC12 cells with a construct containing an EGR promoter linked to SEAP reporter, activation of PC12 cells can be assessed.

The EGR/SEAP reporter construct can be assembled by the following protocol. The EGR-1 promoter sequence (-633 to +1)(Sakamoto K et al., Oncogene 6:867-871 (1991)) can be PCR amplified from human genomic DNA using the following primers:

- 5' GCGCTCGAGGGATGACAGCGATAGAACCCCGG -3' (SEQ ID NO:6) 5' GCGAAGCTTCGCGACTCCCCGGATCCGCCTC-3' (SEQ ID NO:7)
- Using the GAS:SEAP/Neo vector produced in Example 12, EGR1 amplified product can then be inserted into this vector. Linearize the GAS:SEAP/Neo vector using restriction enzymes XhoI/HindIII, removing the GAS/SV40 stuffer. Restrict the EGR1 amplified product with these same enzymes. Ligate the vector and the EGR1 promoter.

To prepare 96 well-plates for cell culture, two mls of a coating solution (1:30 dilution of collagen type I (Upstate Biotech Inc. Cat#08-115) in 30% ethanol (filter sterilized)) is added per one 10 cm plate or 50 ml per well of the 96-well plate, and allowed to air dry for 2 hr.

PC12 cells are routinely grown in RPMI-1640 medium (Bio Whittaker) containing 10% horse serum (JRH BIOSCIENCES, Cat. # 12449-78P), 5% heat-inactivated fetal bovine serum (FBS) supplemented with 100 units/ml penicillin and 100 ug/ml streptomycin on a precoated 10 cm tissue culture dish. One to four split is done every three to four days. Cells are removed from the plates by scraping and resuspended with pipetting up and down for more than 15 times.

Transfect the EGR/SEAP/Neo construct into PC12 using the Lipofectamine protocol described in Example 11. EGR-SEAP/PC12 stable cells are obtained by growing the cells in 300 ug/ml G418. The G418-free medium is used for routine

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growth but every one to two months, the cells should be re-grown in 300 ug/ml G418 for couple of passages.

To assay for neuronal activity, a 10 cm plate with cells around 70 to 80% confluent is screened by removing the old medium. Wash the cells once with PBS (Phosphate buffered saline). Then starve the cells in low serum medium (RPMI-1640 containing 1% horse serum and 0.5% FBS with antibiotics) overnight.

The next morning, remove the medium and wash the cells with PBS. Scrape off the cells from the plate, suspend the cells well in 2 ml low serum medium. Count the cell number and add more low serum medium to reach final cell density as $5x10^5$ cells/ml.

Add 200 ul of the cell suspension to each well of 96-well plate (equivalent to $1x10^5$ cells/well). Add 50 ul supernatant produced by Example 11, 37°C for 48 to 72 hr. As a positive control, a growth factor known to activate PC12 cells through EGR can be used, such as 50 ng/ul of Neuronal Growth Factor (NGF). Over fifty-fold induction of SEAP is typically seen in the positive control wells. SEAP assay the supernatant according to Example 17.

Example 16: High-Throughput Screening Assay for T-cell Activity

NF-κB (Nuclear Factor κB) is a transcription factor activated by a wide variety of agents including the inflammatory cytokines IL-1 and TNF, CD30 and CD40, lymphotoxin-alpha and lymphotoxin-beta, by exposure to LPS or thrombin, and by expression of certain viral gene products. As a transcription factor, NF-κB regulates the expression of genes involved in immune cell activation, control of apoptosis (NF-κB appears to shield cells from apoptosis), B and T-cell development, anti-viral and antimicrobial responses, and multiple stress responses.

In non-stimulated conditions, NF- κB is retained in the cytoplasm with I- κB (Inhibitor κB). However, upon stimulation, I- κB is phosphorylated and degraded, causing NF- κB to shuttle to the nucleus, thereby activating transcription of target genes. Target genes activated by NF- κB include IL-2, IL-6, GM-CSF, ICAM-1 and class 1 MHC.

Due to its central role and ability to respond to a range of stimuli, reporter constructs utilizing the NF-kB promoter element are used to screen the supernatants produced in Example 11. Activators or inhibitors of NF-kB would be useful in treating

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diseases. For example, inhibitors of NF-kB could be used to treat those diseases related to the acute or chronic activation of NF-kB, such as rheumatoid arthritis.

To construct a vector containing the NF-κB promoter element, a PCR based strategy is employed. The upstream primer contains four tandem copies of the NF-κB binding site (GGGGACTTTCCC) (SEQ ID NO:8), 18 bp of sequence complementary to the 5' end of the SV40 early promoter sequence, and is flanked with an XhoI site: 5':GCGGCCTCGAGGGGACTTTCCCGGGGACTTTCCGGGACTTTCCGGGACTTTCCGGGACTTTCCGGGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGGAGACTTTCCGAGACTTTCCGGAGACTTTCCGAGACTTTCCGAGACTTTCCGAGACTTTCCGGAGACTTTCCGAGACTTTCCGAGACTTTCCGAGACTTTCAGACTTCAGACTTTCAGACTTTC

The downstream primer is complementary to the 3' end of the SV40 promoter and is flanked with a Hind III site:

5':GCGGCAAGCTTTTTGCAAAGCCTAGGC:3' (SEQ ID NO:4)

PCR amplification is performed using the SV40 promoter template present in the pB-gal:promoter plasmid obtained from Clontech. The resulting PCR fragment is digested with XhoI and Hind III and subcloned into BLSK2-. (Stratagene) Sequencing with the T7 and T3 primers confirms the insert contains the following sequence:

5':CTCGAGGGGACTTTCCCGGGGACTTTCCGGGGACTTTCC ATCTGCCATCTCAATTAGTCAGCAACCATAGTCCCGCCCCTAACTCCGCCCA TCCCGCCCCTAACTCCGCCCAGTTCCGCCCATTCTCCGCCCCATGGCTGACT AATTTTTTTATTTATGCAGAGGCCGAGGCCGCCTCGGCCTCTGAGCTATTC CAGAAGTAGTGAGGAGGCTTTTTTGGAGGCCTAGGCTTTTGCAAAAAAGCTT: 3' (SEQ ID NO:10)

Next, replace the SV40 minimal promoter element present in the pSEAP2-promoter plasmid (Clontech) with this NF-κB/SV40 fragment using XhoI and HindIII. However, this vector does not contain a neomycin resistance gene, and therefore, is not preferred for mammalian expression systems.

In order to generate stable mammalian cell lines, the NF-κB/SV40/SEAP

cassette is removed from the above NF-κB/SEAP vector using restriction enzymes SalI and NotI, and inserted into a vector containing neomycin resistance. Particularly, the NF-κB/SV40/SEAP cassette was inserted into pGFP-1 (Clontech), replacing the GFP gene, after restricting pGFP-1 with SalI and NotI.

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Once NF-kB/SV40/SEAP/Neo vector is created, stable Jurkat T-cells are created and maintained according to the protocol described in Example 13. Similarly, the method for assaying supernatants with these stable Jurkat T-cells is also described in Example 13. As a positive control, exogenous TNF alpha (0.1,1, 10 ng) is added to wells H9, H10, and H11, with a 5-10 fold activation typically observed.

Example 17: Assay for SEAP Activity

As a reporter molecule for the assays described in Examples 13-16, SEAP activity is assayed using the Tropix Phospho-light Kit (Cat. BP-400) according to the following general procedure. The Tropix Phospho-light Kit supplies the Dilution, Assay, and Reaction Buffers used below.

Prime a dispenser with the 2.5x Dilution Buffer and dispense 15 μ l of 2.5x dilution buffer into Optiplates containing 35 μ l of a supernatant. Seal the plates with a plastic sealer and incubate at 65°C for 30 min. Separate the Optiplates to avoid uneven heating.

Cool the samples to room temperature for 15 minutes. Empty the dispenser and prime with the Assay Buffer. Add 50 μ l Assay Buffer and incubate at room temperature 5 min. Empty the dispenser and prime with the Reaction Buffer (see the table below). Add 50 μ l Reaction Buffer and incubate at room temperature for 20 minutes. Since the intensity of the chemiluminescent signal is time dependent, and it takes about 10 minutes to read 5 plates on luminometer, one should treat 5 plates at each time and start the second set 10 minutes later.

Read the relative light unit in the luminometer. Set H12 as blank, and print the results. An increase in chemiluminescence indicates reporter activity.

Reaction Buffer Formulation:

Reaction Burier Formulation.						
# of plates	Rxn buffer diluent (ml)	CSPD (ml)				
10	60	3				
11	65	3.25				
12	70	3.5				
13	75	3.75				
14	80	4				
15	85	4.25				
16	90	4.5				
17	95	4.75				
18	100	5				
19	105	5.25				
20	110	5.5				
21	115	5.75				
22	120	6				

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23	125	6.25
24	130	6.5
25	135	6.75
26	140	7
27	145	7.25
28	150	7.5
29	155	7.75
30	160	8
31	165	8.25
32	170	8.5
33	175	8.75
34	180	9
35	185	9.25
36	190	9.5
37	195	9.75
38	200	10
39	205	10.25
40	210	10.5
41	215	10.75
42	220	11
43	225	11.25
44	230	11.5
45	235	11.75
46	240	12
47	245	12.25
48	250	12.5
49	255	12.75
50	260	13

Example 18: High-Throughput Screening Assay Identifying Changes in Small Molecule Concentration and Membrane Permeability

Binding of a ligand to a receptor is known to alter intracellular levels of small molecules, such as calcium, potassium, sodium, and pH, as well as alter membrane potential. These alterations can be measured in an assay to identify supernatants which bind to receptors of a particular cell. Although the following protocol describes an assay for calcium, this protocol can easily be modified to detect changes in potassium, sodium, pH, membrane potential, or any other small molecule which is detectable by a fluorescent probe.

The following assay uses Fluorometric Imaging Plate Reader ("FLIPR") to measure changes in fluorescent molecules (Molecular Probes) that bind small molecules. Clearly, any fluorescent molecule detecting a small molecule can be used instead of the calcium fluorescent molecule, fluo-3, used here.

For adherent cells, seed the cells at 10,000-20,000 cells/well in a Co-star black 96-well plate with clear bottom. The plate is incubated in a $\rm CO_2$ incubator for 20 hours. The adherent cells are washed two times in Biotek washer with 200 ul of HBSS (Hank's Balanced Salt Solution) leaving 100 ul of buffer after the final wash.

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A stock solution of 1 mg/ml fluo-3 is made in 10% pluronic acid DMSO. To load the cells with fluo-3, 50 ul of 12 ug/ml fluo-3 is added to each well. The plate is incubated at 37° C in a CO_2 incubator for 60 min. The plate is washed four times in the Biotek washer with HBSS leaving 100 ul of buffer.

For non-adherent cells, the cells are spun down from culture media. Cells are re-suspended to 2-5x10⁶ cells/ml with HBSS in a 50-ml conical tube. 4 ul of 1 mg/ml fluo-3 solution in 10% pluronic acid DMSO is added to each ml of cell suspension. The tube is then placed in a 37°C water bath for 30-60 min. The cells are washed twice with HBSS, resuspended to 1x10⁶ cells/ml, and dispensed into a microplate, 100 ul/well. The plate is centrifuged at 1000 rpm for 5 min. The plate is then washed once in Denley CellWash with 200 ul, followed by an aspiration step to 100 ul final volume.

For a non-cell based assay, each well contains a fluorescent molecule, such as fluo-3. The supernatant is added to the well, and a change in fluorescence is detected.

To measure the fluorescence of intracellular calcium, the FLIPR is set for the following parameters: (1) System gain is 300-800 mW; (2) Exposure time is 0.4 second; (3) Camera F/stop is F/2; (4) Excitation is 488 nm; (5) Emission is 530 nm; and (6) Sample addition is 50 ul. Increased emission at 530 nm indicates an extracellular signaling event which has resulted in an increase in the intracellular Ca⁺⁺ concentration.

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Example 19: High-Throughput Screening Assay Identifying Tyrosine Kinase Activity

The Protein Tyrosine Kinases (PTK) represent a diverse group of transmembrane and cytoplasmic kinases. Within the Receptor Protein Tyrosine Kinase RPTK) group are receptors for a range of mitogenic and metabolic growth factors including the PDGF, FGF, EGF, NGF, HGF and Insulin receptor subfamilies. In addition there are a large family of RPTKs for which the corresponding ligand is unknown. Ligands for RPTKs include mainly secreted small proteins, but also membrane-bound and extracellular matrix proteins.

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Activation of RPTK by ligands involves ligand-mediated receptor dimerization, resulting in transphosphorylation of the receptor subunits and activation of the cytoplasmic tyrosine kinases. The cytoplasmic tyrosine kinases include receptor associated tyrosine kinases of the src-family (e.g., src, yes, lck, lyn, fyn) and non-receptor linked and cytosolic protein tyrosine kinases, such as the Jak family, members of which mediate signal transduction triggered by the cytokine superfamily of receptors (e.g., the Interleukins, Interferons, GM-CSF, and Leptin).

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Because of the wide range of known factors capable of stimulating tyrosine kinase activity, the identification of novel human secreted proteins capable of activating tyrosine kinase signal transduction pathways are of interest. Therefore, the following protocol is designed to identify those novel human secreted proteins capable of activating the tyrosine kinase signal transduction pathways.

Seed target cells (e.g., primary keratinocytes) at a density of approximately 25,000 cells per well in a 96 well Loprodyne Silent Screen Plates purchased from Nalge Nunc (Naperville, IL). The plates are sterilized with two 30 minute rinses with 100% ethanol, rinsed with water and dried overnight. Some plates are coated for 2 hr with 100 ml of cell culture grade type I collagen (50 mg/ml), gelatin (2%) or polylysine (50 mg/ml), all of which can be purchased from Sigma Chemicals (St. Louis, MO) or 10% Matrigel purchased from Becton Dickinson (Bedford,MA), or calf serum, rinsed with PBS and stored at 4°C. Cell growth on these plates is assayed by seeding 5,000 cells/well in growth medium and indirect quantitation of cell number through use of alamarBlue as described by the manufacturer Alamar Biosciences, Inc. (Sacramento, CA) after 48 hr. Falcon plate covers #3071 from Becton Dickinson (Bedford,MA) are used to cover the Loprodyne Silent Screen Plates. Falcon Microtest III cell culture plates can also be used in some proliferation experiments.

To prepare extracts, A431 cells are seeded onto the nylon membranes of Loprodyne plates (20,000/200ml/well) and cultured overnight in complete medium. Cells are quiesced by incubation in serum-free basal medium for 24 hr. After 5-20 minutes treatment with EGF (60ng/ml) or 50 ul of the supernatant produced in Example 11, the medium was removed and 100 ml of extraction buffer ((20 mM HEPES pH 7.5, 0.15 M NaCl, 1% Triton X-100, 0.1% SDS, 2 mM Na3VO4, 2 mM Na4P2O7 and a cocktail of protease inhibitors (# 1836170) obtained from Boeheringer Mannheim (Indianapolis, IN) is added to each well and the plate is shaken on a rotating shaker for 5 minutes at 4°C. The plate is then placed in a vacuum transfer manifold and the extract filtered through the 0.45 mm membrane bottoms of each well using house vacuum. Extracts are collected in a 96-well catch/assay plate in the bottom of the vacuum manifold and immediately placed on ice. To obtain extracts clarified by centrifugation, the content of each well, after detergent solubilization for 5 minutes, is removed and centrifuged for 15 minutes at 4°C at 16,000 x g.

Test the filtered extracts for levels of tyrosine kinase activity. Although many methods of detecting tyrosine kinase activity are known, one method is described here.

Generally, the tyrosine kinase activity of a supernatant is evaluated by determining its ability to phosphorylate a tyrosine residue on a specific substrate (a

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biotinylated peptide). Biotinylated peptides that can be used for this purpose include PSK1 (corresponding to amino acids 6-20 of the cell division kinase cdc2-p34) and PSK2 (corresponding to amino acids 1-17 of gastrin). Both peptides are substrates for a range of tyrosine kinases and are available from Boehringer Mannheim.

The tyrosine kinase reaction is set up by adding the following components in order. First, add 10ul of 5uM Biotinylated Peptide, then 10ul ATP/Mg₂₊ (5mM ATP/50mM MgCl₂), then 10ul of 5x Assay Buffer (40mM imidazole hydrochloride, pH7.3, 40 mM beta-glycerophosphate, 1mM EGTA, 100mM MgCl₂, 5 mM MnCl₂, 0.5 mg/ml BSA), then 5ul of Sodium Vanadate(1mM), and then 5ul of water. Mix the components gently and preincubate the reaction mix at 30°C for 2 min. Initial the reaction by adding 10ul of the control enzyme or the filtered supernatant.

The tyrosine kinase assay reaction is then terminated by adding 10 ul of 120mm EDTA and place the reactions on ice.

Tyrosine kinase activity is determined by transferring 50 ul aliquot of reaction mixture to a microtiter plate (MTP) module and incubating at 37°C for 20 min. This allows the streptavadin coated 96 well plate to associate with the biotinylated peptide. Wash the MTP module with 300ul/well of PBS four times. Next add 75 ul of anti-phospotyrosine antibody conjugated to horse radish peroxidase(anti-P-Tyr-POD(0.5u/ml)) to each well and incubate at 37°C for one hour. Wash the well as above.

Next add 100ul of peroxidase substrate solution (Boehringer Mannheim) and incubate at room temperature for at least 5 mins (up to 30 min). Measure the absorbance of the sample at 405 nm by using ELISA reader. The level of bound peroxidase activity is quantitated using an ELISA reader and reflects the level of tyrosine kinase activity.

Example 20: High-Throughput Screening Assay Identifying Phosphorylation Activity

As a potential alternative and/or compliment to the assay of protein tyrosine kinase activity described in Example 19, an assay which detects activation (phosphorylation) of major intracellular signal transduction intermediates can also be used. For example, as described below one particular assay can detect tyrosine phosphorylation of the Erk-1 and Erk-2 kinases. However, phosphorylation of other molecules, such as Raf, JNK, p38 MAP, Map kinase kinase (MEK), MEK kinase, Src, Muscle specific kinase (MuSK), IRAK, Tec, and Janus, as well as any other

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phosphoserine, phosphotyrosine, or phosphothreonine molecule, can be detected by substituting these molecules for Erk-1 or Erk-2 in the following assay.

Specifically, assay plates are made by coating the wells of a 96-well ELISA plate with 0.1ml of protein G (1ug/ml) for 2 hr at room temp, (RT). The plates are then rinsed with PBS and blocked with 3% BSA/PBS for 1 hr at RT. The protein G plates are then treated with 2 commercial monoclonal antibodies (100ng/well) against Erk-1 and Erk-2 (1 hr at RT) (Santa Cruz Biotechnology). (To detect other molecules, this step can easily be modified by substituting a monoclonal antibody detecting any of the above described molecules.) After 3-5 rinses with PBS, the plates are stored at 4°C until use.

A431 cells are seeded at 20,000/well in a 96-well Loprodyne filterplate and cultured overnight in growth medium. The cells are then starved for 48 hr in basal medium (DMEM) and then treated with EGF (6ng/well) or 50 ul of the supernatants obtained in Example 11 for 5-20 minutes. The cells are then solubilized and extracts filtered directly into the assay plate.

After incubation with the extract for 1 hr at RT, the wells are again rinsed. As a positive control, a commercial preparation of MAP kinase (10ng/well) is used in place of A431 extract. Plates are then treated with a commercial polyclonal (rabbit) antibody (1ug/ml) which specifically recognizes the phosphorylated epitope of the Erk-1 and Erk-2 kinases (1 hr at RT). This antibody is biotinylated by standard procedures. The bound polyclonal antibody is then quantitated by successive incubations with Europium-streptavidin and Europium fluorescence enhancing reagent in the Wallac DELFIA instrument (time-resolved fluorescence). An increased fluorescent signal over background indicates a phosphorylation.

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Example 21: Method of Determining Alterations in a Gene Corresponding to a Polynucleotide

RNA isolated from entire families or individual patients presenting with a phenotype of interest (such as a disease) is be isolated. cDNA is then generated from these RNA samples using protocols known in the art. (See, Sambrook.) The cDNA is then used as a template for PCR, employing primers surrounding regions of interest in SEQ ID NO:X. Suggested PCR conditions consist of 35 cycles at 95°C for 30 seconds; 60-120 seconds at 52-58°C; and 60-120 seconds at 70°C, using buffer solutions described in Sidransky, D., et al., Science 252:706 (1991).

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PCR products are then sequenced using primers labeled at their 5' end with T4 polynucleotide kinase, employing SequiTherm Polymerase. (Epicentre Technologies).

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The intron-exon borders of selected exons is also determined and genomic PCR products analyzed to confirm the results. PCR products harboring suspected mutations is then cloned and sequenced to validate the results of the direct sequencing.

PCR products is cloned into T-tailed vectors as described in Holton, T.A. and Graham, M.W., Nucleic Acids Research, 19:1156 (1991) and sequenced with T7 polymerase (United States Biochemical). Affected individuals are identified by mutations not present in unaffected individuals.

Genomic rearrangements are also observed as a method of determining alterations in a gene corresponding to a polynucleotide. Genomic clones isolated according to Example 2 are nick-translated with digoxigenindeoxy-uridine 5'-triphosphate (Boehringer Manheim), and FISH performed as described in Johnson, Cg. et al., Methods Cell Biol. 35:73-99 (1991). Hybridization with the labeled probe is carried out using a vast excess of human cot-1 DNA for specific hybridization to the corresponding genomic locus.

Chromosomes are counterstained with 4,6-diamino-2-phenylidole and propidium iodide, producing a combination of C- and R-bands. Aligned images for precise mapping are obtained using a triple-band filter set (Chroma Technology, Brattleboro, VT) in combination with a cooled charge-coupled device camera (Photometrics, Tucson, AZ) and variable excitation wavelength filters. (Johnson, Cv. et al., Genet. Anal. Tech. Appl., 8:75 (1991).) Image collection, analysis and chromosomal fractional length measurements are performed using the ISee Graphical Program System. (Inovision Corporation, Durham, NC.) Chromosome alterations of the genomic region hybridized by the probe are identified as insertions, deletions, and translocations. These alterations are used as a diagnostic marker for an associated disease.

Example 22: Method of Detecting Abnormal Levels of a Polypeptide in a Biological Sample

A polypeptide of the present invention can be detected in a biological sample, and if an increased or decreased level of the polypeptide is detected, this polypeptide is a marker for a particular phenotype. Methods of detection are numerous, and thus, it is understood that one skilled in the art can modify the following assay to fit their particular needs.

For example, antibody-sandwich ELISAs are used to detect polypeptides in a sample, preferably a biological sample. Wells of a microtiter plate are coated with specific antibodies, at a final concentration of 0.2 to 10 ug/ml. The antibodies are either monoclonal or polyclonal and are produced by the method described in Example 10.

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The wells are blocked so that non-specific binding of the polypeptide to the well is reduced.

The coated wells are then incubated for > 2 hours at RT with a sample containing the polypeptide. Preferably, serial dilutions of the sample should be used to validate results. The plates are then washed three times with deionized or distilled water to remove unbounded polypeptide.

Next, 50 ul of specific antibody-alkaline phosphatase conjugate, at a concentration of 25-400 ng, is added and incubated for 2 hours at room temperature. The plates are again washed three times with deionized or distilled water to remove unbounded conjugate.

Add 75 ul of 4-methylumbelliferyl phosphate (MUP) or p-nitrophenyl phosphate (NPP) substrate solution to each well and incubate 1 hour at room temperature. Measure the reaction by a microtiter plate reader. Prepare a standard curve, using serial dilutions of a control sample, and plot polypeptide concentration on the X-axis (log scale) and fluorescence or absorbance of the Y-axis (linear scale). Interpolate the concentration of the polypeptide in the sample using the standard curve.

Example 23: Formulating a Polypeptide

The secreted polypeptide composition will be formulated and dosed in a fashion consistent with good medical practice, taking into account the clinical condition of the individual patient (especially the side effects of treatment with the secreted polypeptide alone), the site of delivery, the method of administration, the scheduling of administration, and other factors known to practitioners. The "effective amount" for purposes herein is thus determined by such considerations.

As a general proposition, the total pharmaceutically effective amount of secreted polypeptide administered parenterally per dose will be in the range of about 1 μ g/kg/day to 10 mg/kg/day of patient body weight, although, as noted above, this will be subject to therapeutic discretion. More preferably, this dose is at least 0.01 mg/kg/day, and most preferably for humans between about 0.01 and 1 mg/kg/day for the hormone. If given continuously, the secreted polypeptide is typically administered at a dose rate of about 1 μ g/kg/hour to about 50 μ g/kg/hour, either by 1-4 injections per day or by continuous subcutaneous infusions, for example, using a mini-pump. An intravenous bag solution may also be employed. The length of treatment needed to observe changes and the interval following treatment for responses to occur appears to vary depending on the desired effect.

Pharmaceutical compositions containing the secreted protein of the invention are administered orally, rectally, parenterally, intracistemally, intravaginally,

intraperitoneally, topically (as by powders, ointments, gels, drops or transdermal patch), bucally, or as an oral or nasal spray. "Pharmaceutically acceptable carrier" refers to a non-toxic solid, semisolid or liquid filler, diluent, encapsulating material or formulation auxiliary of any type. The term "parenteral" as used herein refers to modes of administration which include intravenous, intramuscular, intraperitoneal, intrasternal, subcutaneous and intraarticular injection and infusion.

The secreted polypeptide is also suitably administered by sustained-release systems. Suitable examples of sustained-release compositions include semi-permeable polymer matrices in the form of shaped articles, e.g., films, or mirocapsules. 10 Sustained-release matrices include polylactides (U.S. Pat. No. 3,773,919, EP 58,481), copolymers of L-glutamic acid and gamma-ethyl-L-glutamate (Sidman, U. et al., Biopolymers 22:547-556 (1983)), poly (2- hydroxyethyl methacrylate) (R. Langer et al., J. Biomed. Mater. Res. 15:167-277 (1981), and R. Langer, Chem. Tech. 12:98-105 (1982)), ethylene vinyl acetate (R. Langer et al.) or poly-D- (-)-3-hydroxybutyric 15 acid (EP 133,988). Sustained-release compositions also include liposomally entrapped polypeptides. Liposomes containing the secreted polypeptide are prepared by methods known per se: DE 3,218,121; Epstein et al., Proc. Natl. Acad. Sci. USA 82:3688-3692 (1985); Hwang et al., Proc. Natl. Acad. Sci. USA 77:4030-4034 (1980); EP 52,322; EP 36,676; EP 88,046; EP 143,949; EP 142,641; Japanese Pat. Appl. 83-118008; U.S. Pat. Nos. 4,485,045 and 4,544,545; and EP 102,324. Ordinarily, the liposomes 20 are of the small (about 200-800 Angstroms) unilamellar type in which the lipid content is greater than about 30 mol. percent cholesterol, the selected proportion being adjusted for the optimal secreted polypeptide therapy.

For parenteral administration, in one embodiment, the secreted polypeptide is formulated generally by mixing it at the desired degree of purity, in a unit dosage injectable form (solution, suspension, or emulsion), with a pharmaceutically acceptable carrier, i.e., one that is non-toxic to recipients at the dosages and concentrations employed and is compatible with other ingredients of the formulation. For example, the formulation preferably does not include oxidizing agents and other compounds that are known to be deleterious to polypeptides.

Generally, the formulations are prepared by contacting the polypeptide uniformly and intimately with liquid carriers or finely divided solid carriers or both. Then, if necessary, the product is shaped into the desired formulation. Preferably the carrier is a parenteral carrier, more preferably a solution that is isotonic with the blood of the recipient. Examples of such carrier vehicles include water, saline, Ringer's solution, and dextrose solution. Non-aqueous vehicles such as fixed oils and ethyl oleate are also useful herein, as well as liposomes.

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The carrier suitably contains minor amounts of additives such as substances that enhance isotonicity and chemical stability. Such materials are non-toxic to recipients at the dosages and concentrations employed, and include buffers such as phosphate, citrate, succinate, acetic acid, and other organic acids or their salts; antioxidants such as ascorbic acid; low molecular weight (less than about ten residues) polypeptides, e.g., polyarginine or tripeptides; proteins, such as serum albumin, gelatin, or immunoglobulins; hydrophilic polymers such as polyvinylpyrrolidone; amino acids, such as glycine, glutamic acid, aspartic acid, or arginine; monosaccharides, disaccharides, and other carbohydrates including cellulose or its derivatives, glucose, manose, or dextrins; chelating agents such as EDTA; sugar alcohols such as mannitol or sorbitol; counterions such as sodium; and/or nonionic surfactants such as polysorbates, poloxamers, or PEG.

The secreted polypeptide is typically formulated in such vehicles at a concentration of about 0.1 mg/ml to 100 mg/ml, preferably 1-10 mg/ml, at a pH of about 3 to 8. It will be understood that the use of certain of the foregoing excipients, carriers, or stabilizers will result in the formation of polypeptide salts.

Any polypeptide to be used for therapeutic administration can be sterile. Sterility is readily accomplished by filtration through sterile filtration membranes (e.g., 0.2 micron membranes). Therapeutic polypeptide compositions generally are placed into a container having a sterile access port, for example, an intravenous solution bag or vial having a stopper pierceable by a hypodermic injection needle.

Polypeptides ordinarily will be stored in unit or multi-dose containers, for example, sealed ampoules or vials, as an aqueous solution or as a lyophilized formulation for reconstitution. As an example of a lyophilized formulation, 10-ml vials are filled with 5 ml of sterile-filtered 1% (w/v) aqueous polypeptide solution, and the resulting mixture is lyophilized. The infusion solution is prepared by reconstituting the lyophilized polypeptide using bacteriostatic Water-for-Injection.

The invention also provides a pharmaceutical pack or kit comprising one or more containers filled with one or more of the ingredients of the pharmaceutical compositions of the invention. Associated with such container(s) can be a notice in the form prescribed by a governmental agency regulating the manufacture, use or sale of pharmaceuticals or biological products, which notice reflects approval by the agency of manufacture, use or sale for human administration. In addition, the polypeptides of the present invention may be employed in conjunction with other therapeutic compounds.

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Example 24: Method of Treating Decreased Levels of the Polypeptide

It will be appreciated that conditions caused by a decrease in the standard or normal expression level of a secreted protein in an individual can be treated by administering the polypeptide of the present invention, preferably in the secreted form. Thus, the invention also provides a method of treatment of an individual in need of an increased level of the polypeptide comprising administering to such an individual a pharmaceutical composition comprising an amount of the polypeptide to increase the activity level of the polypeptide in such an individual.

For example, a patient with decreased levels of a polypeptide receives a daily dose 0.1-100 ug/kg of the polypeptide for six consecutive days. Preferably, the polypeptide is in the secreted form. The exact details of the dosing scheme, based on administration and formulation, are provided in Example 23.

Example 25: Method of Treating Increased Levels of the Polypeptide

Antisense technology is used to inhibit production of a polypeptide of the present invention. This technology is one example of a method of decreasing levels of a polypeptide, preferably a secreted form, due to a variety of etiologies, such as cancer.

For example, a patient diagnosed with abnormally increased levels of a polypeptide is administered intravenously antisense polynucleotides at 0.5, 1.0, 1.5, 2.0 and 3.0 mg/kg day for 21 days. This treatment is repeated after a 7-day rest period if the treatment was well tolerated. The formulation of the antisense polynucleotide is provided in Example 23.

Example 26: Method of Treatment Using Gene Therapy

One method of gene therapy transplants fibroblasts, which are capable of expressing a polypeptide, onto a patient. Generally, fibroblasts are obtained from a subject by skin biopsy. The resulting tissue is placed in tissue-culture medium and separated into small pieces. Small chunks of the tissue are placed on a wet surface of a tissue culture flask, approximately ten pieces are placed in each flask. The flask is turned upside down, closed tight and left at room temperature over night. After 24 hours at room temperature, the flask is inverted and the chunks of tissue remain fixed to the bottom of the flask and fresh media (e.g., Ham's F12 media, with 10% FBS, penicillin and streptomycin) is added. The flasks are then incubated at 37°C for approximately one week.

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At this time, fresh media is added and subsequently changed every several days. After an additional two weeks in culture, a monolayer of fibroblasts emerge. The monolayer is trypsinized and scaled into larger flasks.

pMV-7 (Kirschmeier, P.T. et al., DNA, 7:219-25 (1988)), flanked by the long terminal repeats of the Moloney murine sarcoma virus, is digested with EcoRI and HindIII and subsequently treated with calf intestinal phosphatase. The linear vector is fractionated on agarose gel and purified, using glass beads.

The cDNA encoding a polypeptide of the present invention can be amplified using PCR primers which correspond to the 5' and 3' end sequences respectively as set forth in Example 1. Preferably, the 5' primer contains an EcoRI site and the 3' primer includes a HindIII site. Equal quantities of the Moloney murine sarcoma virus linear backbone and the amplified EcoRI and HindIII fragment are added together, in the presence of T4 DNA ligase. The resulting mixture is maintained under conditions appropriate for ligation of the two fragments. The ligation mixture is then used to transform bacteria HB101, which are then plated onto agar containing kanamycin for the purpose of confirming that the vector has the gene of interest properly inserted.

The amphotropic pA317 or GP+am12 packaging cells are grown in tissue culture to confluent density in Dulbecco's Modified Eagles Medium (DMEM) with 10% calf serum (CS), penicillin and streptomycin. The MSV vector containing the gene is then added to the media and the packaging cells transduced with the vector. The packaging cells now produce infectious viral particles containing the gene (the packaging cells are now referred to as producer cells).

Fresh media is added to the transduced producer cells, and subsequently, the media is harvested from a 10 cm plate of confluent producer cells. The spent media, containing the infectious viral particles, is filtered through a millipore filter to remove detached producer cells and this media is then used to infect fibroblast cells. Media is removed from a sub-confluent plate of fibroblasts and quickly replaced with the media from the producer cells. This media is removed and replaced with fresh media. If the titer of virus is high, then virtually all fibroblasts will be infected and no selection is required. If the titer is very low, then it is necessary to use a retroviral vector that has a selectable marker, such as neo or his. Once the fibroblasts have been efficiently infected, the fibroblasts are analyzed to determine whether protein is produced.

The engineered fibroblasts are then transplanted onto the host, either alone or after having been grown to confluence on cytodex 3 microcarrier beads.

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It will be clear that the invention may be practiced otherwise than as particularly described in the foregoing description and examples. Numerous modifications and variations of the present invention are possible in light of the above teachings and, therefore, are within the scope of the appended claims.

The entire disclosure of each document cited (including patents, patent applications, journal articles, abstracts, laboratory manuals, books, or other disclosures) in the Background of the Invention, Detailed Description, and Examples is hereby incorporated herein by reference.

	(1) GENERAL INFORMATION:
5	(i) APPLICANT: Human Genome Sciences, Inc. et al.(ii) TITLE OF INVENTION: 87 Human Secreted Proteins(iii) NUMBER OF SEQUENCES: 323(iv) CORRESPONDENCE ADDRESS:
10	 (A) ADDRESSEE: Human Genome Sciences, Inc. (B) STREET: 9410 Key West Avenue (C) CITY: Rockville (D) STATE: Maryland (E) COUNTRY: USA (F) ZIP: 20850
15	(v) COMPUTER READABLE FORM:
20	(A) MEDIUM TYPE: Diskette, 3.50 inch, 1.4Mb storage(B) COMPUTER: HP Vectra 486/33(C) OPERATING SYSTEM: MSDOS version 6.2(D) SOFTWARE: ASCII Text
25	(vi) CURRENT APPLICATION DATA:(A) APPLICATION NUMBER:(B) FILING DATE: March 19, 1998(C) CLASSIFICATION:
30	<pre>(vii) PRIOR APPLICATION DATA: (A) APPLICATION NUMBER: (B) FILING DATE:</pre>
35	<pre>(viii) ATTORNEY/AGENT INFORMATION: (A) NAME: A. Anders Brookes (B) REGISTRATION NUMBER: 36,373 (C) REFERENCE/DOCKET NUMBER: PZ004PCT</pre>
40	(vi) TELECOMMUNICATION INFORMATION: (A) TELEPHONE: (301) 309-8504 (B) TELEFAX: (301) 309-8439
45	(2) INFORMATION FOR SEQ ID NO: 1:
50	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 733 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear
55	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 1:

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GGGATCCGGA GCCCAAATCT TCTGACAAAA CTCACACATG CCCACCGTGC CCAGCACCTG

	AATTCGAGGG TGCACCGTCA GTCTTCCTCT TCCCCCCAAA ACCCAAGGAC ACCCTCATGA	120
5	TCTCCCGGAC TCCTGAGGTC ACATGCGTGG TGGTGGACGT AAGCCACGAA GACCCTGAGG	180
	TCAAGTTCAA CTGGTACGTG GACGGCGTGG AGGTGCATAA TGCCAAGACA AAGCCGCGGG	240
10	AGGAGCAGTA CAACAGCACG TACCGTGTGG TCAGCGTCCTT CACCGTCCTG CACCAGGACT	300
10	GGCTGAATGG CAAGGAGTAC AAGTGCAAGG TCTCCAACAA AGCCCTCCCA ACCCCCATCG	360
	AGAAAACCAT CTCCAAAGCC AAAGGGCAGC CCCGAGAACC ACAGGTGTAC ACCCTGCCCC	420
15	CATCCCGGGA TGAGCTGACC AAGAACCAGG TCAGCCTGAC CTGCCTGGTC AAAGGCTTCT	480
	ATCCAAGCGA CATCGCCGTG GAGTGGGAGA GCAATGGGCA GCCGGAGAAC AACTACAAGA	540
20	CCACGCCTCC CGTGCTGGAC TCCGACGGCT CCTTCTTCCT CTACAGCAAG CTCACCGTGG	600
20	ACAAGAGCAG GTGGCAGCAG GGGAACGTCT TCTCATGCTC CGTGATGCAT GAGGCTCTGC	660
	ACAACCACTA CACGCAGAAG AGCCTCTCCC TGTCTCCGGG TAAATGAGTG CGACGGCCGC	720
25	GACTCTAGAG GAT	733
30	(2) INFORMATION FOR SEQ ID NO: 2:	
	() OPOUTNOT OUR DECEMBRATION	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 5 amino acids	
35	(B) TYPE: amino acid (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 2:	
	Trp Ser Xaa Trp Ser	
40	1 5	
45	(2) INFORMATION FOR SEO ID NO: 3:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 86 base pairs	
50	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
50	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 3:	
55	GCGCCTCGAG ATTTCCCCGA AATCTAGATT TCCCCGAAAT GATTTCCCCG AAATGATTTC	60
	CCCGAAATAT CTGCCATCTC AATTAG	86

	(2) INFORMATION FOR SEQ ID NO: 4:	
5	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 27 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 4:	
	GCGGCAAGCT TTTTGCAAAG CCTAGGC	27
15		
15	(2) INFORMATION FOR SEQ ID NO: 5:	
20	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 271 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 5:	
	CTCGAGATTT CCCCGAAATC TAGATTTCCC CGAAATGATT TCCCCGAAAT GATTTCCCCG	60
••	AAATATCTGC CATCTCAATT AGTCAGCAAC CATAGTCCCG CCCCTAACTC CGCCCATCCC	120
30	GCCCCTAACT CCGCCCAGTT CCGCCCATTC TCCGCCCCAT GGCTGACTAA TTTTTTTTAT	180
	TTATGCAGAG GCCGAGGCCG CCTCGGCCTC TGAGCTATTC CAGAAGTAGT GAGGAGGCTT	240
35	TTTTGGAGGC CTAGGCTTTT GCAAAAAGCT T	271
40	(2) INFORMATION FOR SEQ ID NO: 6:	
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 32 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 6:	
50	GCGCTCGAGG GATGACAGCG ATAGAACCCC GG	32
55	(2) INFORMATION FOR SEQ ID NO: 7:	
	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 31 base pairs(B) TYPE: nucleic acid	
60	(C) STRANDEDNESS: double	

	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 7:	
5	GCGAAGCTTC GCGACTCCCC GGATCCGCCT C	31
10	(2) INFORMATION FOR SEQ ID NO: 8:	
15	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 12 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 8:	12
25		
23	(2) INFORMATION FOR SEQ ID NO: 9:	
30	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 73 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 9:	
	GCGGCCTCGA GGGGACTTTC CCGGGGACTT TCCGGGGACT TTCCGGGACT TTCCATCCTG	60
40	CCATCTCAAT TAG	73
	(2) INFORMATION FOR SEQ ID NO: 10:	
45	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 256 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double	
50	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 10:	
	CTCGAGGGGA CTTTCCCGGG GACTTTCCGG GGACTTTCCA TCTGCCATCT	60
55	CAATTAGTCA GCAACCATAG TCCCGCCCCT AACTCCGCCC ATCCCGCCCC TAACTCCGCC	120
	CAGTTCCGCC CATTCTCCGC CCCATGGCTG ACTAATTTTT TTTATTTATG CAGAGGCCGA	180
60	GGCCGCCTCG GCCTCTGAGC TATTCCAGAA GTAGTGAGGA GGCTTTTTTG GAGGCCTAGG	240

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	25
CTTTTGCAAA AAGCTT	• • • • • • • • • • • • • • • • • • • •

5 (2) INFORMATION FOR SEQ ID NO: 11:

10

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1679 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 11:

	(X1) SEQUENCE DESCRIPTION. SEQ 15 No. 11.	
15	GCAGCGCACC CGGGCGATCG CTTCACGGAT GCGGACGACG TAGCCATCCT TACCTACGTG	60
	AAGGAAAATG CCCGCTCGCC CAGCTCCGTC ACCGGTAACG CCTTGTGGAA AGCGATGGAG	120
20	AAGAGCTCGC TCACGCAGCA CTCGTGGCAG TCCCTGAAGG ACCGCTACCT CAAGCACCTG	180
	CGGGGCCAGG AGCATAAGTA CCTGCTGGGG GACGCGCCGG TGAGCCCCTC CTCCCAGAAG	240
25	CTCAAGCGGA AGGCGGAGGA GGACCCGGAG GCCGCGGATA GCGGGGAACC ACAGAATAAG	300
25	AGAACTCCAG ATTTGCCTGA AGAAGAGTAT GTGAAGGAAG AAATCCAGGA GAATGAAGAA	360
	GCAGTCAAAA AGATGCTTGT GGAAGCCACC CGGGAGTTTG AGGAGGTTGT GGTGGATGAG	420
30	AGCCCTCCTG ATTTTGAAAT ACATATAACT ATGTGTGATG ATGATCCACC CACACCTGAG	480
	GAAGACTCAG AAACACAGCC TGATGAGGAG GAAGAAGAAG AAGAAGAAAA AGTTTCTCAA	540
25	CCAGAGGTGG GAGCTGCCAT TAAGATCATT CGGCAGTTAA TGGAGAAGTT TAACTTGGAT	600
35	CTATCAACAG TTACACAGGC CTTCCTAAAA AATAGTGGTG AGCTGGAGGC TACTTCCGCC	660
	TTCTTAGCGT CTGGTCAGAG AGCTGATGGA TATCCCATTT GGTCCCGACA AGATGACATA	720
40	GATTTGCAAA AAGATGATGA GGATACCAGA GAGGCATTGG TCAAAAAATT TGGTGCTCAG	780
	AATGTAGCTC GGAGGATTGA ATTTCGAAAG AAATAATTGG CAAGATAATG AGAAAAAGAAA	840
45	AAAGTCATGG TAGGTGAGGT GGTTAAAAAA AATTGTGACC AATGAACTTT AGAGAGTTCT	900
43	TGCATTGGAA CTGGCACTTA TTTTCTGACC ATCGCTGCTG TTGCTCTGTG AGTCCTAGAT	960
	TTTTGTAGCC AAGCAGAGTT GTAGAGGGGG ATAAAAAGAA AAGAAATTGG ATGTATTTAC	1020
50	AGCTGTCCTT GAACAAGTAT CAATGTGTTT ATGAAAGGAA GATCTAAATC AGACAGGAGT	1080
	TGGTCTACAT AGTAGTAATC CATTGTTGGA ATGGAACCCT TGCTATAGTA GTGACAAAGT	1140
55	GAAAGGAAAT TTAGGAGGCA TAGGCCATTT CAGGCAGCAT AAGTAATCTC CTGTCCTTTG	1200
33	GCAGAAGCTC CTITAGATTG GGATAGATTC CAAATAAAGA ATCTAGAAAT AGGAGAAGAT	1260
	TTAATTATGA GGCCTTGAAC ACGGATTATC CCCAAACCCT TGTCATTTCC CCCAGTGAGC	1320
60	TCTGATTTCT AGACTGCTTT GAAAATGCTG TATTCATTTT GCTAACTTAG TATTTGGGTA	1380

	CCCTGCTCTT TGGCTGTTCT TTTTTTGGAG CCCTTCTCAG TCAAGTCTGC CGGATGTCTT	1440
5	TCTTTACCTA CCCCTCAGTT TTCCTTAAAA CGCGCACACA ACTCTAGAGA GTGTTAAGAA	1500
J	TAATGTTACT TGGTTAATGT GTTATTTATT GAGTATTGTT TGTGCTAAGC ATTGTGTTAG	1560
	ATTTAAAAAA TTAGTGGATT GACTCCACTT TGTTGTTGTTG TTTTCATTGT TGAAAATAAA	1620
10	TATAACTTTG TATTCGAAAA AAAAAAAAA AAAATNRCTG CGGNCCGACA AGGGAATTC	1679
15	(2) INFORMATION FOR SEQ ID NO: 12:	
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 1830 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 12:	
25	GCGACCGCGC CCTTCAGCTA GCTCGCTCGC TCGCTCTGCT TCCCTGCTGC CGGCTGCGCA	60
	TGGCTTNGGC GTTGGCGGCG CTGGCGGCGG TCGAGCNGCC TGCGSAGCCG GTACCAGCAG	120
30	TTGCAGAATG AAGAAGAGTC TGGAGAACCT GAACAGGCTG CAGGTGATGC TCCTCCACCT	180
20	TACAGCAGCA TTTCTGCAGA GAGCGCACAT NATTTTGACT ACAAGGATGA GTCTGGGTTT	240
	CCAAAGCCCC CATCTTACAA TGTAGCTACA ACACTGCCCA GTTATGATGA AGCGGAGAGG	300
35	ACCAAGGCTG AAGCTACTAT CCCTTTGGTT CCTGGGAGAG ATGAGGATTT TGTGGGTCGG	360
	GATGATTTTG ATGATGCTGA CCAGCTGAGG ATAGGAAATG ATGGGATTTT CATGTTAACT	420
40	TTTTTCATGG CATTCCTCTT TAACTGGATT GGGTTTTTCC TGTCTTTTTG CCTGACCACT	480
	TCAGCTGCAG GAAGGTATGG GGCCATTTCA GGATTTGGTC TCTCTCTAAT TAAATGGATC	540
	CTGATTGTCA GGTTTTCCAC CTATTTCCCT GGATATTTTG ATGGTCAGTA CTGGCTCTGG	600
45	TGGGTGTTCC TTGTTTTAGG CTTTCTCCTG TTTCTCAGAG GATTTATCAA TTATGCAAAA	660
	GTTCGGAAGA TGCCAGAAAC TTTCTCAAAT CTCCCCAGGA CCAGAGTTCT CTTTATTTAT	720
50	TAAAGATGTT TTCTGGCAAA GGCCTTCCTG CATTTATGAA TTCTCTCTCA AGAAGCAAGA	780
	GAACACCTGC AGGAAGTGAA TCAAGATGCA GAACACAGAG GAATAATCAC CTGCTTTAAA	840
	AAAATAAAGT ACTGTTGAAA AGATCATTTC TCTCTATTTG TTCCTAGGTG TAAAATTTTA	900
55	ATAGTTAATG CAGAATTCTG TAATCATTGA ATCATTAGTG GTTAATGTTT GAAAAAGCTC	960
	TTGCAATCAA GTCTGTGATG TATTAATAAT GCCTTATATA TTGTTTGTAG TCATTTTAAG	1020
60	TAGCATGAGC CATGTCCCTG TAGTCGGTAG GGGGCAGTCT TGCTTTATTC ATCCTCCATC	1080

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	TCAAAATGAA	CTTGGAATTA	AATATTGTAA	GATATGTATA	ATGCTGGCCA	TTTTAAAGGG	1140
	GTTTTCTCAA	AAGTTAAACT	TTTGTTATGA	CTGTGTTTTT	GCACATAATC	CATATTTGCT	1200
5	GTTCAAGTTA	ATCTAGAAAT	TTATTCAATT	CTGTATGAAC	ACCTGGAAGC	AAAATCATAG	1260
	TGCAAAAATA	CATTTAAGGT	GTGGTCAAAA	ATAAGTCTTT	AATTGGTAAA	TAATAAGCAT	1320
	TAATTTTTTA	TAGCCTGTAT	TCACAATTCT	GCGGTACCTT	ATTGTACCTA	AGGGATTCTA	1380
10	AAGGTGTTGT	CACTGTATAA	AACAGAAAGC	ACTAGGATAC	AAATGAAGCT	TAATTACTAA	1440
	AATGTAATTC	TTGACACTCT	TTCTATAATT	AGCGTTCTTC	ACCCCCACCC	CCACCCCCAC	1500
15	CCCCCTTATT	TTCCTTTTGT	CTCCTGGTGA	TTAGGCCAAA	GTCTGGGAGT	AAGGAGAGGA	1560
	TTAGGTACTT	AGGAGCAAAG	AAAGAAGTAG	CTTGGAACTT	TTGAGATGAT	CCCTAACATA	1620
	CTGTACTACT	TGCTTTTACA	ATGTGTTAGC	AGAAACCAGT	GGGTTATAAT	GTAGAATGAT	1680
20	GTGCTTTCTG	CCCAAGTGGT	AATTCATCTT	GGTTTGCTAT	GTTAAAACTG	TAAATACAAC	1740
	AGAACATTAA	TAAATATCTC	TTGTGTAGCA	CCTTTTAAAA	AAAAAAAAA	AAAAAAAA	1800
25	AAAAAAAA	AANCCCGGGG	GGGGGCCCCI	ī			1830

30 (2) INFORMATION FOR SEQ ID NO: 13:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1212 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 13:

TGTTGAAGT TGTTACTTTT GTTTACAGCA AAGTTTGATG TAGTGTGCAG TAGTGAGCTC 60 40 TAGACTGATC TTTTTCTAAA TCAGAAAGTG ATTAAAGTAT GCACAACCAA AGGCAGGTTT 120 TTCTTTTTCA TTTATTCAGC AACTATTTAT TAAGCATCAA CTCTGTGCCA GGCACGTTAC 180 45 TAGCTGCTAC ATACTGTCTG AACATGACAT ACGGTTAAGT AACTTTACAA TTATTATCAA 240 ATACTTCAAT GTAGATATTT CTTAAGTTGA AATAGCATTA ACTAGGATAA TGCTTTCATG 300 360 TTATTTATT TGTCTTGTGA TAGAAATTCA ACTTTGTACC ATCTTAAAAC TAGGTTGCTA 50 TAAAAATAGG AGGATGAAGT CAATAAAGTT TATGCCAGTT TAAAAAACTGG AAGGAAAAGG 420 TAAGAGCTCT CCATTATAAA ATAGTTGCAT TCGGTTAATT TTTACACATT AGTGCATTGC 480 55 GTATATCAAC TGGCCCTCAA TGAAGCATTT AAGTGCTTGG AATTTTACTA AACTGACTTT 540 TTTGCAACTT TGGGAGATTT TTGAGGGGAG TGTTGAAAAAT TGCCAAACAC TCACCTCTTA 600 CTCAAAACTT CAAATAAAAT ACACATTTC AAGAGGGAGC ACCTTTTATA TTTGATAAGT 660 60

	TTTCATTATA AACCTTATAA TACCAGTCAC AAAGAGGTTG TCTGTCTATG GTTTAGCAAA	720
5	CATTTGCTTT TCTTTTTGGA AGTGTGATTG CAATTGCAGA ACAGAAAGTG AGAAAACACT	780
J	GCCAGCGGTG ATTGCTACTT GAGGTAGTTT TTTACAACTA CCATTTCCCC TCCATGAAAT	840
	TATGTGAAAT TTATTTTATC TTTGGGAAAA GTTGAGAAGA TAGTAAAAGA ATTAGGAATT	900
10	TAAAATTACA GGGAAAAATA TGTAAGTGAA AAGCAATAAA TATTTTGTTC ACTTTGCTAT	960
	CAAGATGTTC ACTATCAGAT ATTTATTATA TGGCAGCAAT TTATATTTTT AATCATTGCC	1020
15	CATTAATAGA CGCAGTAAAA TATTTTTGAA TCAGACATTT GGGGTTTGTA TGTGCATTAA	1080
13	AATTGTCTTT TGTACTGTAA GTTACTGTTA ATTTGAATAT TTTATTGAAC TGTCTCCCTG	1140
	TGCCTTTATA ATATAAAGTT GTTTCTACAA CTTTTAATGA TCTTAATAAA GAATACTTTA	1200
20	AGAAAAAA AA	1212
25	(2) INTERPORTATION FOR CEO ID NO. 14	
23	(2) INFORMATION FOR SEQ ID NO: 14:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 2061 base pairs	
30	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
	(D) TOPOLOGY: linear	
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 14:	
35	GGTTTTCCTC CGACTTCCGG ACATCTCCCT GGGAGTCGCG CAGAGTGGAG TCAAAGGCAA	60
	CCAGTGCTCG CTGCGGTCTC TGGGGATCGG GACCGCGGCG GCGGCCCGCG AGCGGGATGT	120
40	TCCGGGGCTT GAGCAGTTGG TTGGGCTTGC AGCAGCCGGT GGCAGCCCA	180
	ATGGAGATGC TCCACCCGAG CAGCCGTCCG AGACGGTGGC TGAGTCTGCG GAGGAGGAGC	240
	TGCAGCAAGC GGGAGACCAG GAGCTCCTCC ACCAGGCCAA AGACTTCGGC AACTATTTAT	300
45	TTAACTTTGC ATCTGCTGCC ACAAAAAAGA TAACTGAATC AGTTGCTGAA ACAGCACAAA	360
	CAATAAAGAA ATCCGTAGAA GAAGGAAAAA TAGATGGCAT CATTGACAAG ACAATTATAG	420
50	GAGATTTTCA GAAGGAACAG AAAAAATTTG TTGAAGAGCA ACATACAAAG AAGTCAGAAG	480
	CAGCTGTGCC CCCATGGGTT GACACTAACG ATGAAGAAAC AATTCAACAA CAAATTTTGG	540
	CCTTATCAGC TGACAAGAGG AATTTCCTTC GTGACCCTCC GGCTGGCGTG CAATTTAATT	600
55	TCGACTTTGA TCAGATGTAC CCCGTGGCCC TGGTCATGCT CCAGGAGGAT GAGCTGCTAR	660
	CAAGATGAGA TTTGCCCTCG TTCCTAAACT TGTGAAGGAA GAAGTGTTCT GGAGGAACTA	720
60	CTTTTACCGC GTCTCCCTGA TTAAGCAGTC AGCCCAGCTC ACGGCCCTGG CTGCCCAACA	780

	GCAGGCCGCA GGGAAGGGAG GAGAAGAGCA ATGGCAGAGA GCAAGATTTG CCGCTGGAGA	840
-	GGCAGTACGG CCCAAAACGC CACCCGTTGT AATCAAATCT CAGCTTAAAA CTCAAGAGGA	900
5	TGAGGAAGAA ATTTCTACTA GCCCAGGTGT TTCTGAGTTT GTCAGTGATG CCTTCGATGC	960
	CTGTAACCTA AATCAGGAAG ATCTAAGGAA AGAAATGGAG CAACTAGTGC TTGACAAAAA	1020
10	GCAAGAGGAG ACAGCCGTAC TGGAAGAGGA TTCTGCAGAT TGGGAAAAAG AACTGCAGCA	1080
10	GGAACTTCAA GAATATGAAG TGGTGACAGA ATCTGAAAAA CGAGATGAAA ACTGGGATAA	1140
	GGAAATAGAG AAAATGCTTC AAGAGGAAAA TTAGCTGTTC CTGAAATAGA AGAATAATCC	1200
15	TTAACAGTCT GCAAACTGAC ATTAAATTCT AGATGTTGAC AATTACTGAA TCAGAAGGCA	1260
	TGAAAGAGTA TAATTTTATG AAATTCAAAA TTATTCTTTT TTCAAGTTGA AACTTGCCTC	1320
20	TTCTACTTTA AAAAAGTATA TAGAACAGTT ACTTCTAATA ATCAGAAAGA GATGTTTTAT	1380
20	AGAACATTTC TTTAATATAA AGTTAGAGAT GTCTTCATAG GCAGTATGGC TATCTTTGCC	1440
	ACAGAAACAT AAGTAAAATT TTAGAGTTCT GTTTTCCATG AGGTCAAAAA TATAATTTAT	1500
25	TCCTCAGTCA TGGTTTTCTA AATATCTGTA CTCCACATTC CATTTTAATT GATATGAGGG	1560
	TGTTAAAGTA CCTACTTAAT GGGTTGATTA CTATCAAAAT GACCAAATTA TACCAAAGAA	1620
20	CTTAAGAGGA AGCACTTTCA GAACTATTCA CTTGCCAGGT ATTTTCTAAA ATTCCACCTG	1680
30	AAAGCCAAAA GATAAAATAC ATNAGTTGGA TTTTAATGAT ATAAGCATCA CACAATTTTA	1740
	CATTAAGAAA TACTGTGCAG CCCATGCGTG GTGGCTCAGG CCTGTAATCC CAGCANTTTG	1800
35	GGAGGCCGAG GTGGGCAGAT CACCGGAGGT CAGGAGTTCG AGACCAGCCT TGCCAACATA	1860
	GTGAAACCCT GTCTTTACTA AAAATACAAA AATTAGCCGG GCATGGTGGC AGGCACCTGT	1920
40	AATCCCAGCT ACTAGGGAGG CTTTTGAACC CAGGAGGCAG AGGTTGCAGC GAGCTGAGAT	1980
40	CGCGCCACTG CACTCCAGCC TGGGTGATAG AGTGAGATTC AGTCTCAAAA AAAAAAAAAA	2040
	AAAAAAAA AATGACCTCG A	2061
45		
	(2) INFORMATION FOR SEQ ID NO: 15:	
50	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1412 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
55	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 15:	
	CCCTTCATCT GCGTTGCCAG GAACCCTGTC AGCAGAAACT TCTCAAGCCC CATCCTTGCC	60

AGGAAGCTCT GTGAAGGTGC TGCTGATGAC CCAGATTCCT CCATGGTCCT CCTGTGTCTC

	CTGTTGGTGC	CCCTCCTGCT	CAGTCTCTTT	GTACTGGGGC	TATTTCTTTG	GTTTCTGAAG	180
5	AGAGAGAGAC	AAGAAGAGTA	CATTGAAGAG	AAGAAGAGAG	TGGACATTTG	TCGGGAAACT	240
3	CCTAACATAT	GCCCCCATTC	TGGAGAGAAC	ACAGAGTACG	ACACAATCCC	TCACACTAAT	300
	AGAACAATCC	TAAAGGAAGA	TCCAGCAAAT	ACGGTTTACT	CCACTGTGGA	AATACCGAAA	360
10	AAGATGGAAA	ATCCCCACTC	ACTGCTCACG	ATGCCAGACA	CACCAAGGCT	ATTIGCCTAT	420
	GAGAATGTTA	TCTAGACAGC	AGTGCACTCC	CCTAAGTCTC	TGCTCAAAAA	AAAAACAATT	480
15	CTCGGCCCAA	AGAAAACAAT	CAGAAGAATT	CACTGATTTG	ACTAGAAACA	TCAAGGAAGA	540
1.5	ATGAAGAACG	TTGACTTTTT	TCCAGGATAA	ATTATCTCTG	ATGCTTCTTT	AGATTTAAGA	600
	GTTCATAATT	CCATCCACTG	CTGAGAAATC	TCCTCAAACC	CAGAAGGTTT	AATCACTTCA	660
20	TCCCAAAAAT	GGGATTGTGA	ATGTCAGCAA	ACCATAAAAA	AAGTGCTTAG	AAGTATTCCT	720
	ATAAAAATGT	AAATGCAAGG	TCACACATAT	TAATGACAGC	CTGTTGTATT	AATGATGGCT	780
25	CCAGGTCAGT	GTCTGGAGTT	TCATTCCATC	CCAGGGCTTG	GATGTCAGGA	TTATACCAAG	840
	AGTCTTGCTA	CCAGGAGGC	AAGAAGACCA	AAACAGACAG	ACAAGTCCAG	CAGAAGCAGA	900
	TGCACCTGAC	AAAAATGGAT	GTATTAATTG	GCTCTATAAA	CTATGTGCCC	AGCAYTATGC	960
30	TGAGCTTACA	CTAATTGGTC	AGACATGCTG	TCTGCCCTCA	TGAAATTGGC	TCCAAATGAW	1020
	TGAACTACTT	TCATGAGCAG	TTGTAGCAGG	CCTGACCACA	GATTCCCAGA	GGGCCAGGTG	1080
35	TGGATCCACA	GGACTTGAAG	GTCAAAGTTC	ACAAAGATGA	AGAATCAGGG	TAGCTGACCA	1140
	TGTTTGGCAG	ATACTATAAT	GGAGACACAG	AAGTGTGCAT	GGCCCAAGGA	CAAGGACCTC	1200
	CAGCCAGGCT	TCATTTATGC	ACTTGTCTGC	AAAAGAAAAG	TCTAGGTTTT	AAGGCTGTGC	1260
40	CAGAACCCAT	CCCAATAAAG	AGACCGAGTC	TGAAGTCACA	TTGTAAATCT	AGTGTAGGAG	1320
	ACTTGGAGTC	AGGCAGTGAG	ACTGGTGGG	CACGGGGGGC	ANTGGGTANT	GTAAACCTTT	1380
1 5	TAAAGATGGT	TAATTCNTCA	TTAGTGTTTT	TT			1412

(2) INFORMATION FOR SEQ ID NO: 16:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1052 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 16:

TTCCTCTCT CTCTCTACCC CTCCTGTCTC TCCTCCCCTC CTCTCTTCTTC CTCTCCTCTC 60

420

	TCTCTTCCTC TCCTCTCT TCCCTTCCTG TCTCTCTTCC CCTCCTCTCT CTCTTCCTGT	120
	CCTCTATCTC TTCCCCTCCT CTATCTCTTC CTCTCCTCTC TCTCTTCCTC TCCTCTCTCT	180
5	CTCTTSCTTT CTTCTCTCTC TCCTGTCTCG GCTGTTGTGG GTTGCAGGTT GGGTGCTGCT	240
	GTTGTGGTCC TTCCCAGAAA CTGCCAGTAG AGGGCAGCCT GGGCATCCTA ATGCTTACTC	300
	TGGTTGTTAC ACAAAGAAAA TATTGGGGTC ACTGGCGAGC CCACCCACAC TCACCAGAAT	360
l O.	CTCCACTGTA GTCCCCCTAA CAAACAGCCC TTCACTTCCT CTCCCACTTC AGCAATTTGT	420
	ATTTTGATGC CATTGGCCTC AGATCAGAGT GTTTTAAATC ATCACGCCCT GGCTTATCCC	480
15	TGGTCGAGCC AGGACACGGG GTGCTTCAGT GGGTCTGTCA CCCTCTCTCC TTGAAGCATG	540
	TTGCTTTTAT TTATTTACTT TTACTCTCAC CCTGCTCCTG TACCAGCAGG GGCCACTTCA	600
	AAGCCAAGGT ACAGGGTGAT AACTTGTGGT CCAGCATCAG TTTTCTCCAC TTCTTTCTCC	660
20	CACTCACCCC CAGCAAGGTG CCTGGGGAGA CTTGAGCAGA TGTTTCATTT TGGCCTGGCC	720
	AGTGGCTGAA AGCAGGCCTC CAATGCACTG TGACCTCTGG CTTCCCCAGC AGCTTTCCCA	780
25	GAGAGGCAGA GGGGCCTTCC ACAGCCCGGG TTCTCCTGCT GCCTCCTGCC TGCTGCAGCT	840
	GCAGGCATTC TGAGGGGCAA CGTGGAGGAA GGGCCAGGGA TGCATGGGAT TTTAATTGTT	900
•	TCATCACACC TTCCCCGTGG CAAAGAAACA GTCAGTCCTC TTCAGGTGTC TTCTGGATTT	960
30	CTGGTGATGG ACAGAGAAAT CITTTTACAG TTTCAAATTA TGTTCAACAA ATAAAAATTG	1020
	CATTTTTAT TTTGGAAAAA AAAAAAAAAA AA	1052
35		
	TO THE PROPERTY OF THE PROPERT	
	(2) INFORMATION FOR SEQ ID NO: 17:	
40	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 683 base pairs	
	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
45	(D) TOPOLOGY: linear	
45	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 17:	
	AATTCGGCAG AGGCACTTAT CATGTACATA TAGCCTGTTT TITAGCATTG TTAGACAAAG	60
50	TAGGCATATT CCTTTCCATC CAAGAACTCA TAACCTAGTA ATTGTAGTTG GCTGATAGCT	120
	CATTGCCCAT ACACAAGGAT CTAACACAAC CTCTTGAATA AACATCCCCC TTATTCAGAA	180
==	ATGCCTTTTC CTATTTCCAT ATTGCAACTT TGCTTACAAA TTTCCAATCT GTCTTTCTGT	240
55	TTACAGAAGA TATACAAAAT TCCTTTTGTA TGATCTCTTT ATATCTCTTG ATTTTCTTTT	300

GTGTTTGCTA CCAAAGGGCC TGCACATAGT GAGAAGATTG TGCATGATCT GTGAGCTCTA

CCACACCTGG AATTAGGGAT CACCAATATG AGAAAAAAAA TTGGAGGTAC AAATAACATT

	ATCATATGTW	ATTGGCATAT	AAATTACAGA	TGTWTCTATG	ACTAAAAACC	CTGTGGATAT	480
5	WAACCMAATG	CAGATAAWTW	TAATAAATW	TWTAAAAATW	TWATCMAATA	ATGATAGTGC	540
3	TATTCAAATA	CTTCAAATTT	GCACAGTGAT	TTATTTCTTA	AAATATGTTA	ACACATGTGA	600
	GCCAATACAC	TGAGGTCACT	GGATAAATAA	ACAGATTCTT	GCAAAAAAA	AAAAAAAA	660
10	ACTCGAGGGG	GGCCCGTACC	CTT				683
15	(2) INFORMA	ATION FOR SE	EQ ID NO: 18	3:			
			HARACTERIST				
20	, =,	(A) LENG (B) TYP: (C) STR	GTH: 1054 b E: nucleic ANDEDNESS: OLOGY: line	ase pairs acid double			
	(xi)	SEQUENCE I	DESCRIPTION	: SEQ ID NO	: 18:		
25	AAACTCATTT	AGGTGACACT	ATAGAAGGTA	CGCCTGCAGG	TACCGGTCCG	GAATTCCCGG	60
	GTCGACCCAC	GMGNCCGGCG	ACAAGATGGC	AGCAGCGTGT	CGGAGCGTGA	AGGGCCTGGT	120
30	GGCGGTAATA	ACCGGAGGAG	CCTCGGGCCT	GGGCCTGGCC	ACGGCGGACG	ACTTGTGGGG	180
30	CAGGGAGCCT	CTGCTGTGCT	TCTGGACCTG	CCCAACTCGG	GTGGGGAGGC	CCAAGCCAAG	240
	AAGTTAGGAA	ACAACTGCGT	TTTCGCCCCA	GCCGACGTGA	CCTCTGAGAA	GGATGTGCAA	300
35	ACAGCTCTGG	CTCTAGCAAA	AGGAAAGTTT	GCCGTGTGG	ATGTAGCTGT	CAACTGTGCA	360
	GGCATCGCGG	TGGCTAGCAA	GACGTACAAC	TTAAAGAAGG	GCCAGACCCA	TACCTTGGAA	420
40	GACTTCCAGC	GAGTTCTTGA	TGTGAATCTC	ATGGGCACCT	TCAATGTGAT	CCGCCTGGTG	480
	GCTGGTGAGA	TGGGCCAGAA	TGAACCAGAC	CAGGGAGGCC	AACGTGGGGT	CATCATCAAC	540
	ACTGCCAGTG	TGGCTGCCTT	CGAGGGTCAG	GTTGGACAAG	CTGCATACTC	TGCTTCCAAG	600
45	GGGGAATAG	TGGGCATGAC	ACTGCCCATT	GCTCGGGATC	TGGCTCCCAT	AGGTATCCGG	660
	GTGATGACCA	TTGCCCCAGG	TCTGTTTGGC	ACCCCACTGC	TGACCAGCCT	CCCAGAGAAA	720
50	GTGTGCAACT	TCTTGGCCAG	CCAAGTGCCC	TTCCCTAGCC	GACTGGGTGA	CCCTGCTGAG	780
	TATGCTCACC	TCGTACAGGC	CATCATCGAG	AACCCATTCC	TCAATGGAGA	GGTCATCCGG	840
	CTGGATGGG	CCATTCGTAT	GCAGCCTTGA	AGGGAGAAGG	CAGAGAAAAC	ACACGCTCCT	900
55	CTGCCCTTCC	TTTCCCTGGG	GTACTACTCT	CCAGCTTGGG	AGGAAGCCCA	GTAGCCATTT	960
	TGTAACTGCC	TACCAGTCGC	CCTCTGTGCC	TAATAAAGTC	TCTTTTTCTC	ACANAAAAAA	1020
60	АААААААА	АААААААА	АААААААА	AAAA			1054

_	(2) INFORMATION FOR SEQ ID NO: 19:						
5	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1393 base pairs						
10	(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear						
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 19:						
15	GGAACAAGCT GGGATATGTG AGCGTTAAGC TACTCACATC CTTCAAAAAAG GTGAAACATC	60					
15	TTACACGGGA CTGGAGAACC ACAGCACATG CTTTGAAGTA TTCAGTGGTC CTTGAGTTGA	120					
	ATGAGGNCCA CCGGAAGGTG AGGAGGACCA CCCCCGTCCC ACTGTTCCCC AACGAGAACC	180					
20	TCCCCAGCAA GATGCTCCTG GTCTATGATC TCTACTTGTY TCCTAAGCTG TGGGCTCTGG	240					
	CCACCCCCA GAAGAATGGG AAGGGTGCAA GARAAGGTGA TGGAACACCT GCTCAAGCTT	300					
25	TTTGGGACTT TTGGAGTCAT CTCATCAGTG CGGATCCTCA AACCTGGGAG AGAGCTGCCC	360					
25	CCTGACATCC GGAGGNTCCA GCAGCCGCTA CAGCTCCTCT GACCCCGAGA GCAACCCCAC	420					
	ATCCCCTATG GCGGCCCGAC GGCACGNGKC CACCAACAAG CTCAGCCCGT CTGGCCACCA	480					
30	GAATCTCTTT CTGAGTCCAA ATGCCTCCCC GTGCACAAGT CCTTGGAGCA GCCCCTTGGC	540					
	CCAACGCAAA GGCGTTTCCA GAAAGTCCCC ACTGGCGGAG GAAGGTAGAC TGAACTGCAG	600					
25	CACCAGCCCT GAGATCTTCC GCAAGTGTAT GGATTATTCC TCTGACAGCA GCGTCACTCC	660					
35	CTCTGGCAGC CCCTGGGTCC GGAGGCGTCG CCAAGCCGAG ATGGGGGACCC AGGAGAAAAG	720					
	CCCCGGTACG AGTCCCCTGC TCTCCCGGAA GATGCAGACT GCAGATGGGS TACCCGTAGG	780					
40	TNGCTTGAGG TTGCCCAGGG GTCCTGACAA CACCAGAGGA TTTCATGGCC ATGAGAGGAG	840					
	CAGGGCCTGT GTATAAATAC CTTCTATTTT TAATACAAGC TCCACTGAAA ACCACCTTCG	900					
	TTTTCAAGGT TCTGACAAAC ACCTGGCATG ACAGAATGGA ATTCGTTCCC CTTTGAGAGA	960					
45	TTTTTTATTC ATGTAGACCT CTTAATTTAT CTATCTGTAA TATACATAAA TCGGTACGCC	1020					
	ATGGTTTGAA GACCACCTTC TAGTTCAGGA CTCCTGTTCT TCCCAGCATG GCCACTATTT	1080					
50	TGATGATGGC TGATGTGTGT GAGTGTGATG GCCCTGAAGG GCTGTAGGAC GGAGGTTCCC	1140					
	TGGGGGAAGT CTGTTCTTTG GTATGGAATT TTTCTCTCTT CTTTGGTATG GAATTTTTCC	1200					
	CTTCAGTGAC TGAGCTGTCC TCGATAGGCC ATGCAAGGGC TTCCTGAGAG TTCAGGAAAG	1260					
55	TTCTCTTGTG CAACAGCAAG TAGCTAAGCC TATAGCATGG TGTCTTGTAG GACCAAATCG	1320					
	ATGTTACCTG TCAAGTAAAT AAATAATAAA ACACCCAACT GGGAGTGCTG AAAAAAAANA	1380					

60 ANNAAAAAAC TCG

(2) INFORMATION FOR SEQ ID NO: 20:

10	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1215 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 20:	
15	AGGAAAAGTT TTCCNAATTG GAAAGCGGGC AGTGAGCGCA ACGCAATTAA TGTGAGTTAG	60
	NTCANTCATT AGGCACCCCA GGCTTTACAC TTTATGCTTC CGGNTCGTAT GTTGTGTGGA	120
20	ATTGTGAGCG GATAACAATT TCACACAGGA AACAGCTATG ACCATGATTA CGCCAAGCTN	180
20	TAATACGACT CACTATAGGG AAAGCTGGTA CGCCTGCAGG TACCGGTCCG GAATTCCCGG	240
	GTCGACCCAC GCGTCCGCC ACGCGTCCGT GAAAATCCGA AGTGCCGCGG AAAGTGGAGG	300
25	TGAGGGCCGC CCGCCCTAGA GGTGCCCGTC CGAGAGGCAG AGCTGACAAG GAAGGTTTCG	360
	AGCGTTTTGC TGGCAAAGGG ATTTCTTACA ACCTCCAGGC ATGCGTCTTT CTGCCCTGCT	420
30	GGCCTTGGCA TCCAAGGTCA CTCTGCCCCC CCATTACCGC TATGGGATGA GCCCCCCAGG	480
50	CTCTGTTGCA GACAAGAGGA AGAACCCCCC ATGGATCAGG CGGCGCCCAG TGCTTGTGGA	540
	ACCCATCTCT GATGAAGACT GGTATCTGTT CTGTGGGGAC ACGGTGGAGA TC" GAAGG	600
35	CAAGGATGCC GGGAAGCAGG GCAAAGTGGT TCAAGTTATC CGGCAGCGAA ACTUGGTGGT	660
	CGTGGGAGGG CTGAACACAC ATTACCGCTA CATTGGCAAG ACCATGGATT ACCGGGGAAC	720
40	CATGATCCCT AGTGAAGCCC CCTTGCTCCA CCGCCAGGTC AAACTTGTGG ATCCTATGGA	780
10	CAGGAAACCC ACTGAGATCG AGTGGAGATT TACTGAAGCA GGAGAGCGGG TACGAGTCTC	840
	CACACGATCA GGGAGAATTA TCCCTAAACC CGAATTTCCC AGAGCTGATG GCATCGTCCC	900
45	TGAAACGTGG ATTGATGGCC CCAAAGACAC ATCAGTGGAA GATGCTTTAG AAAGAACCTA	960
	TGTGCCCTGT CTAAAGACAC TGCAGGAGGA GGTGATGGAG GCCATGGGGA TCAAGGAGAC	1020
50	CCGGAAATAC AAGAAGGTCT ATTGGTATTG AGCCTGGGGC AGAGCAGCTC CTCCCCAACT	1080
50	TCTGTCCCAG CCTTGAAGGC TGAGGCACTT CTTTTTCAGA TGCCAATAAA GAGCACTTTA	1140
	ТСАСТССТСС АЛАЛАЛАЛА ЛАЛАЛАЛАЛ ЛАЛАЛАЛАЛ ЛАЛАЛАЛА	1200
55	AAAAGGGCCG GCCGC	1215

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(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2042 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double
(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 21:

10	CTGCATCCAG GCGCAGAATA ACCTGGGTAT CTTGTGGTCT GAAAGAGAGA AATTGAAACT	60
	GCACAGGCTT ACCTAGAGTC ATCAGAAGCA CTATATAATC AGTATATGAA AGAGGTTGGG	120
. ~	AGTCCTCCTC TTGATCCTAC TGAGCGTTTT CTTCTGAAGA AGAGAAACTT ACTGAACAAG	180
15	AGAGATCAAA AAGATTTGAA AAGGTTTATA CTCATAACCT ATATTACCTA GCTCAAGTCT	240
	ACCAGCATCT GGAAATGTTT GAGAAGGCTG CTCACTATTG CCATAGTACA CTAAAACGCC	300
20	AGCTTGAGCA CAATGCCTAC CATCCTATAG AGTGGGCTAT CAATGCTGCT ACCTTGTCAC	360
	AGTTTTACAT CAATAAGCTA TGCTTTATGG AGGCCAGGCA CTGTTTATCA GCTGCTAATG	420
25	TCATTITTGG TCAAACTGGA AAGATCTCAG CCACAGAAGA CACTCCTGAA GCTGAAGGAG	480
23	AAGTGCCAGA GCTTTATCAT CAAAGAAAGG GGGAAATAGC AAGGTGCTGG ATCAAATACT	540
	GTTTGACTCT CATGCAGAAT GCCCAACTCT CCATGCAGGA CAACATAGGA GAGCTTGATC	600
30	TTGATAAACA GTCTGAACTT AGAGCTTTAA GGAAAAAAGA ACTAGATGAG GAGGAAAGCA	660
	TTCGGAAAAA AGCTGTGCAG TTTGGAACCG GTGAACTGTG TGATGCCATC TCTGCAGTAG	720
25	AAGAGAAAGT GAGCTACTTG AGACCTTTAG ATTTTGAAGA AGCCAGAGAA CTTTTCTTAT	780
35	TGGGTCAGCA CTATGTCTTT GAGGCAAAAG AGTTCTTTCA GATTGATGGT TATGTCACTG	840
	ACCATATTGA AGTTGTCCAA GACCACAGTG CTCTGTTTAA GGTGCTTGCA TTCTTTGAAA	900
40	CTGACATGGA GAGACGGTGC AAGATGCATA AACGCRGAAT AGCCATGCTA GAGCCCCTAA	960
	CTGTAGACCT GAATCCACAG TATTATCTGT TGGTCAACAG ACAGATCCAG TTTGAAATTG	1020
45	CACATGCTTA CTATGATATG ATGGATTTGA AGGTTGCCAT TGCTGACAGG CTAAGGGATC	1080
73	CTGATTCACA CATTGTAAAA AAAATAAATA ATCTTAATAA GTCAGCACTG AAGTACTACC	1140
	AGCTCTTCTT AGACTCCCTG AGAGACCCAA ATAAAGTATT CCCTGAGCAT ATAGGGGAAG	1200
50	ATGTTCTTCG CCCTGCCATG TTAGCTAAGT TTCGAGTTGC CCGTCTCTAT GGCAAAATCA	1260
	TTACTGCAGA TCCCAAGAAA GAGCTGGAAA ATTTGGCAAC ATCATTGGGA ACATTACAAA	1320
55	TTTATTGTTG ATTACTGTGA AAAGCATCCT GAGGCCGCCC AGGAAATAGA AGTTGAGCTA	1380
55	GAACTTAGTA AAGAGATGGT TAGTCTTCTC CCAACAAAAA TGGAGAGATT CAGAACCAAG	1440
	ATGGCCCTGA CTTAATCCTT GTTTTTAAAG AAAGGAAATG TGCAATATTG AAGTGATCTT	1500
60	TTTCCCTAGT CAGACAGGCC CAATTCCATT GTGATGTTTA CCTTTATAGC CAGGTGAGTG	1560

	CAGTITGAAC	TIGAGATACA	GTCAACTGAG	TGTTTGCTAG	GATCCTAAGG	AACATAAAGT	1620
5	ТААТТАААА	CTTACACCTA	ATTATGTAAA	TTGCCTTGTT	AAAGACATGT	GATTTGTATT	1680
J	TTAGATGCTT	GTTTCCTATT	AAAATACAGA	CATTTCTACC	CTCAGTTTCT	AAATGTAGAC	1740
	TATTTGTTGG	CTAGTACTTG	ATAGATTCCT	TGTAAGAAAA	AATGCTGGGT	AATGTACCTG	1800
10	GTAACAAGCC	TGTTAATATA	TTAAGATTGA	AAAAGTAACT	TCTATAGTTA	CTCCTTCTAA	1860
	AATATTTGAC	TTCCTACATT	CCCCCACCC	AAAATCTTTC	CCTTTTGAAA	ATACTAAAAA	1920
15	CTAAGTTATG	TTATTATAAA	GTGTAAAATG	GTTTGTCTTA	ATTATAGGAG	AAAAAGGCCT	1980
13	TGTTAGAAAT	AAAATAAACT	GACTTATTTC	ACTAATGAAA	AAAAAAAA	AAAAAAAA	2040
	тт						2042
20							
	(2) INFORMA	TION FOR SE	EQ ID NO: 22).			
25		SEQUENCE CH (A) LENG (B) TYPP (C) STR	HARACTERIST GTH: 1872 b E: nucleic ANDEDNESS: OLOGY: line	ICS: ase pairs acid double			
30	(xi)			: SEQ ID NO	: 22:		
	GGGTCGACCC	ACGCGTCCGA	TTGGCCTAGA	GCTCCTGTGA	CCGAGAGCGC	CACGGAAGCC	60
35	TGGGGATGAT	GTCGGGCAGC	TTTATTCTTT	GCTTGGCTTT	GGTAACTAGG	TGGTCCCCTC	120
	AAGCATCCTC	AGTTCCTCTT	GCTGTTTATG	AATCTAAGAC	AAGGAAGTCC	TATAGAAGCC	180
40	AAAGGGACAG	GGACGGAAAG	GACAGGTCCC	AAGGGATGGG	GCTGTCTTTA	CTTGTGGAAA	240
. •	CCAGGAAATT	GCTCCTCTCA	GCCAACCAAG	GTTGACCACA	CACCACCCTT	CCGGAGCAGC	300
	TCAGTCAGCC	CTCGGGGACG	RGAAACCACA	AGCGCAGAGA	CGCTGAGGCC	CAGGCAGGTG	360
45	AAGAGGAAGT	GGCTTTGGGT	TTTTAAAGTA	GGTGAGCGTG	ACCTCTCTGA	CTGCTTCTTC	420
	cccgggggg	ACTGCAAACC	GCTCAGGGTT	GCGGCAGAGC	CATGGACTTC	CGGTCCCTGC	480
50	AACGGGTGAC	CTAAGCGTGG	TGCACCCATC	AGTCACGCAG	GAGGACTGAC	TTGACAGACG	540
	AAAGACAAGC	CCGGATGACA	CAGGGTGAGA	AGAGTCAGGG	CCGCACCTCT	GTCCCTGCAA	600
	ACCAACAGGT	GCATGGTGAG	TGTGGCAGTC	CCCACAGCTC	CACAATGGGC	TCCCCCGCCA	660
55	ACGGGGACGA	CAGGGATCTT	CAGGAACTTC	TGACCTCACC	AAGTCAAGTG	GACCACTCTC	720
	CACTCCACGA	GGATGTGAAA	CGGTTCTTTA	AAATGGGATT	TTAGAGCCTC	GGGAATGCAT	780
60	GTGCGTCGCA	TCTTTCATAT	TATGGGTCAG	GATAGATTCA	TTTCTTGCAA	CATAGTGGAA	840

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	AAGATATAAG	CTGCAGTAAT	TTGCTCTTTG	AATGACCGTC	ACCCCCAGTA	TAGGATATGC	900
	TTGTATCCCC	CCGTCACTCC	TCCGCCTGTT	TTTTAAACTT	TTCCACCACC	TGCGTCCAAA	960
5	AAGAATGTTA	TAGCGAGTGC	TCTTAAATGT	TGAACCTGGG	TGTTGCTTCC	GGGCCAGTCT	1020
	GCGTGGCTCC	ATGAAAAGCT	CACTGCTGCC	CCAGCCGGGC	TTCTTAGAGG	AGGTCAGTTG	1080
10	TCCTATGTAT	CATCATTTAC	TCTGGGAATC	CTACTGTGAA	ATCATGTCTG	TATTTTTCTG	1140
10	GAGCAGTTCA	CATAGAGTAG	AATGTGGAAT	TTCCCGTGAA	CGTCTCCTTC	CTCCCCCGTA	1200
	TCTGCCGCCT	GTCACTTCGC	CACCGTGCTA	GAATACTGTT	GTGTTGTAAG	ATGACTAATT	, 1260
15	TTAAAAGAAC	CTGCCCTGAA	AAGTTCTTAG	AAACGCAATG	AAAGGGAGGA	ACTTGTCCTT	1320
	TACCCAGTTT	TTCCTTTGTA	GGATGGGAAA	GTATAAAAAG	GCACAGAAGG	TTGTCATGGG	1380
	CTGTTCCTTG	GGGGTTTTTA	TCCTGCTCAC	CGTGGAGATA	AGCCTGCGGC	TTGTCTAACC	1440
20	AGCGCAGCGM	AAAGGTCTCA	ATGCCTTTTG	GTAACATCCG	TCATTGCAGA	AGAAAGTTTA	1500
	CACGACGTCA	AAAAGTGACG	TTCATGCTAA	GTGTTTTTCC	AGAAATATTG	GTTTCATGTT	1560
25	TCTTATTKGC	TCTGCCTCCT	GTGCTTATAT	CATCCAAAAA	CTTTTTAAAA	AGGTCCAGAA	1620
	TTCTATTTA	ACCTGATGTT	GAGCACCTTT	AAAACGTTCG	TATGTGTGTT	GCACTAATTC	1680
20	TAAACTTTGG	AGGCATTITG	CTGTGTGAGG	CCGATCGCCA	CTGTAAAGGT	CCTAGAGTTG	1740
30	CCTGTTTGTC	TCTGGAGATG	GAATTAAACC	AAATAAAGAG	CTTCCACTGG	AGGCTTGTAT	1800
	TGACCTTGTA	ACTATATGTT	AATCTCGTGT	TAAAATAAAA	TATAACTTGT	GAAAAAAAAA	1860
35	AAAAAAAA	NT					1872

40 (2) INFORMATION FOR SEQ ID NO: 23:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 289 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 23:

60

(2) INFORMATION	FOR	SEQ	ID	NO:	24
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5	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 3533 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 24:	
	TTTTATTTAC TTCAAATTAA CTGTACTTTA CTCAAATAGA AAANGAATAA TTTTCACATT	60
15	ATGAAGCTAC ACAATTCCAA AATACACATG CTGAGGCTCT TTTTAAGTCC GAATTGTCTA	120
	GTAATTACAA AAAAGTGAAG AGTTTACAGA TATACAAGGA AATAAAGGCG AATTATTGCA	180
20	AAGAAAACAA GTTTAATTTC ACTTTGAATG ACAACGATTT TTCTGGAAAG CAGATACTTC	240
	ACTCCTTTAA GTTTCCACCC AAGCCACAAT AATTTCAAAC GGTCTTGCGG ATGACCCAGC	300
	TGGTCACTCT TGTTTATGTG GGGACTGGAG GTAATGAGAG CCAAAAAAAG TGCTATAAAC	360
25	CTAATTTGGC TAGAGCAAGT TCACACGACA CGACCGTGCT TTAAAAACTT GCTCTCCATT	420
	ATGTACTTCC TTCCATCAGG TTGGGGAAAA AAAAATGGTG GGGATGGTGA GTAAACACAC	480
30	CAGTGGTTTC ATCAGAGGGG AACTCACTAC TCAGGAGGTG ACGGTGACGT GGTGCCGGTC	540
50	CCTGAAGTAC GCGCACAAGC TCCGGAGGTT GCGGGAGCTT CCGCTGCCGC CTGGAGGGAA	600
	GCCGGAGCGA CGGGGGTCAC GGCGGCGGTC AGAGGGTAAA GGTCTTGCTC CCAGCAGCCT	660
35	CCGCGGTGGA TACGTCGCCA TCTTGGATCC GCGGGACAAG AAAATTCATG CGAGGGAGAC	720
	GTGGTGGCG GTCCTTCCTG TGACACGACC CTTGAGTGAC AGTTCTATTT GATTGCCTCC	780
40	GGTACTGTGA GGAAAGGACA CGACTCTATG GTGAGGACTG ATGGACATAC ATTATCTGAG	840
	AAAAGAAACT ACCAGGTGAC AAACAGCATG TTTGGTGCTT CAAGAAAGAA GTTTGTAGAG	900
	GGGGTCGACA GTGACTACCA TGACGAAAAC ATGTACTACA GCCAGTCTTC TATGTTTCCA	960
45	CATCGGTCAG AAAAAGATAT GCTGGCATCA CCATCTACAT CAGGTCAGCT GTCTCAGTTT	1020
	GGGGCAAGTT TATACGGGCA ACAAAGTGCA CTAGGCCTTC CAATGAGGGG GATGAGCAAC	1080
50	AATACCCCTC AGTTAAATCG CAGCTTATCA CAAGGCACTC AGTTACCGAG CCACGTCACG	1140
50	CCAACAACAG GGGTACCAAC AATGTCACTT CACACGCCTC CATCTCCAAG CAGGGGTATT	1200
	TTGCCTATGA ATCCTARGAA TATGATGAAC CACTCCCAGG TTGGTCAGGG CATTGGAATT	1260
55	CCTAGCAGGA CAAATAGCAT GAGCAGTTCA GGGTTAGGTA GCCCCAACAG AAGCTCGCCA	1320
	AGCATAATAT GTATGCCAAA GCAGCAGCCT TCTCGACAGC CTTTTACTGT GAACAGTATG	1380
60	TCTGGATTTG GAATGAACAG GAATCAGGCA TTTGGAATGA ATAACTCCTT ATCAAGTAAC	1440

	ATTTTTAATG GAACAGACGG AAGTGAAAAT GTGACAGGAT TGGACCTTTC AGATTTCCCA	1500
	GCATTAGCAG ACCGAAACAG GAGGGAAGGA AGTGGTAACC CAACTCCATT AATAAACCCC	1560
5	TTGGCTGGAA GAGCTCCTTA TGTTGGAATG GTAACAAAAC CAGCAAATGA ACAATCCCAG	1620
	GACTTCTCAA TACACAATGA AGATTTTCCA GCATTACCAG GCTCCAGCTA TAAAGATCCA	1680
10	ACATCAAGTA ATGATGACAG TAAATCTAAT TTGAATACAT CTGGCAAGAC AACTTCAAGT	1740
10	ACAGATGGAC CCAAATTCCC TGGAGATAAA AGTTCAACAA CACAAAATAA TAACCAGCAG	1800
	AAAAAAGGGA TCCAGGTGTT ACCTGATGGT CGGGTTACTA ACATTCCTCA AGGGATGGTG	1860
15	ACGGACCAAT TTGGAATGAT TGGCCTGTTA ACATTTATCA GGGCAGCAGA GACAGACCCA	1920
	GGAATGGTAC ATCTTGCATT AGGAAGTGAC TTAACAACAT TAGGCCTCAA TCTGAACTCT	1980
20	CCTGAAAATC TCTACCCCAA ATTTGCGTCA CCCTGGGCAT CTTCACCTTG TCGACCTCAA	2040
20	GACATAGACT TCCATGTTCC ATCTGAGTAC TTAACGAACA TTCACATTAG GGATAAGCTG	2100
	GCTGCAATAA AACTTGGCCG ATATGGTGAA GACCTTCTCT TCTATCTCTA TTACATGAAT	2160
25	GGAGGAGACG TATTACAACT TTTAGCTGCA GTGGAGCTTT TTAACCGTGA TTGGAGATAC	2220
	CACAAAGAAG AACGAGTATG GATTACCAGG GCACCAGGCA TGGAGCCAAC AATGAAAAACC	2280
30	AATACCTATG AGAGGGGAAC ATATTACTTC TTTGACTGTC TTAACTGGAG GAAAGTAGCT	2340
50	AAGGAGTTCC ATCTGGAATA TGACAAATTA GAAGAACGGC CTCACCTGCC ATCCACCTTC	2400
	AACTACAACC CTGCTCAGCA AGCCTTCTAA AAAAAAAAA AAAAAAAAA AAAAAAAAA AAAAAGACTT	2460
35	CCCTTTTCTT GGGGTATGGC TGTCTCAGCA CAATACTCAA CATAACTGCA GAACTGATGT	2520
	GGCTCAGGCA CCCTGGTTTT AATTCCTTGA GGATCTGGCA ATTGGCTTAC GCAAAAGGTC	2580
40	ACCATTTGAG GTCCTGCCTT ACTAATTATG TGCTGCCCAA CAACTAAATT TGTAATTTGT	2640
••	TTTTCTCTAG TTTGAGCAGG GTCTGAATTT TTTCATTTAT TTCCTTTTTT GCCAGCAGAC	2700
	AGACTTGAGT CTGTAAAGAC AAGCAAATAC ACTGACAGAA GTTTACCATA GTTTCTAAAA	2760
45	TGTAAAAAG AAAACCCCCA AAAGACTCAA GAAAATTAGA CCACAAATTT TGCATTGTTC	2820
	ATTGTAGCAC TATTGGTAAT AAAATAACAA ATGTTTGTGC ATTTTTATGT GAAGATCCTT	2880
50	CTCGTATTTC ATTTGGAAAG ATGAGCAAGA GGTCTGCTTC CTTCATTTTA CTTCCCCTTC	2940
	TGTTTTGAA AGGCAGTTTC GCCAAGCTTA ATGCAAGAAT ATCTGACTGT TTAGAAGAAA	3000
	GATATTGCCA CAATCTCTGG ATGGTTTTCC AGGGTTGTGT TATTACTGAG CTTCATCTTT	3060
55	CCAGAATGAG CAAAACACTG TCCAGTCTTT GTTACGATTT TGTAATAAAT GTGTACATTT	3120
	TTTTTAAATT TTTGGACATC ACATGAATAA AGGTATGTAT GTACGAATGT GTATATATTA	3180
60	TATATATGAC ATCTATTITG GAAAATGTTT GCCCTGCTGT ACCTCATTTT TAGGAGGTGT	3240

	GCATGGATGC	AATATATGAA	AATGGGACAT	TCTGGAACTG	CTGGTCAGGG	GACTTTGTCG	3300				
	CCCTGTGCAC	TAAAAGGCCC	AGATTTTCAG	CAGCCAAGGA	CATCCATACC	CAAGTGAATG	3360				
5	TGATGGGACT	TAAAAGAAGT	GAACTGAGAC	AATTCACTCT	GGCTGTTTGA	ACAGCAGCGT	3420				
	TTCATAGGAA	GAGAAAAAA	GATCAATCTT	GTATTTTCTG	ACCACATAAA	GGCTTCTTCT	3480				
10	CTTTGTAATA	AAGTAGAAAA	GCTCTCCTCA	AAAAAAAA	AAAAAAACTC	GAG	3533				
	(2) INFORMA	(2) INFORMATION FOR SEQ ID NO: 25:									
15	(i)	SEQUENCE CH	ARACTERIST	ICS:							
			GTH: 1148 b E: nucleic	-							
20			ANDEDNESS: O DLOGY: line								
	(xi) SEQUENCE I	DESCRIPTION	: SEQ ID NO	: 25:						
25	ACCCACGCGT	CCGCAAATTA	TACTTCCTCA	TTCATATTAT	GTTGATACAA	AAGACCTTGG	60				
23	CAGCCATTTC	TCCCAGCAGT	TTTAAAGGAT	GAACATTGGA	TTTCATGCCA	TCCCATAGAA	120				
	AACCTGTTTT	AAAATTTTAG	GGATCTTTAC	TTGGTCATAC	ATGAAAAGTA	CACTGCTTAG	180				
30	AAATTATAGA	CTATTATGAT	CTGTCCACAG	TGCCCATTGT	CACTTCTTTG	TCTCATTTCT	240				
	TCCCTTTGTT	CCTTAGTCAT	CCAAATAAGC	CTGAAAACCA	TAAGAGATAT	TACTTTATTG	300				
35	AATATGGTTG	GCATTAAATT	TAGCATTTCA	TTATCTAACA	AAATTAATAT	AAATTCCAGG	360				
	ACATGGTAAA	ATGTGTTTTA	ATAACCCCCA	GACCCAAATG	AAAATTTCAA	AGTCAATACC	420				
	AGCAGATTCA	TGAAAGTAAA	TTTAGTCCTA	TAATTTTCAG	CTTAATTATA	AACAAAGGAA	480				
40	CAAATAAGTG	GAAGGGCAGC	TATTACCATT	CGCTTAGTCA	AAACATTCGG	TTACTGCCCT	540				
	TTAATACACT	CCTATCATCA	GCACTTCCAC	CATGTATTAC	AAGTCTTGAC	CCATCCCTGT	600				
45	CGTAACTCCA	GTAAAAGTTA	CTGTTACTAG	ATTTTTA	TCAATTAACT	GACAAATAGT	660				
	TTCTTTTTAA	AGTAGTTTCT	TCCATCTTTA	TTCTGACTAG	CTTCCAAAAT	GTGTTCCCTT	720				
	TTTGAATCGA	GGTTTTTTTG	TTTTGTTTTG	TTTTCTGAAA	AAATCATACA	ACTTTGTGCT	780				
50	TCTATTGCTT	TTTTGTGTTT	TGTTAAGCAT	GTCCCTTGGC	CCAAATGGAA	GAGGAAATGT	840				
	TTAATTAATG	CTTTTTAGTT	TAAATAAATT	GAATCATTTA	TAATAATCAG	TGTTAACAAT	900				
55	TTAGTGACCC	TTGGTAGGTT	AAAGGTTGCA	TTATTTATAC	TTGAGATTTT	TTTCCCCTAA	960				
						TCAAGAAACA	1020				
						CTTATATAAA	1080				
60	TAAAATTGGT	GGTACTAATG	TGAAAAAAA	ААААААААА	AACTCGAGGG	GGGCCCGGTA	1140				

	CCCTATTA	1148
5		
_	(2) INFORMATION FOR SEQ ID NO: 26:	
10	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 717 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 26:	
	GGCACGAGCT AGCTGCCGCC ACCCGAACAG CCTGTCCTGG TGCCCCGGCT CCCTGCCCCG	60
	CGCCCAGTCA TGACCCTGCG CCCCTCACTC CTCCCGCTCC ATCTGCTGCT GCTGCTGCTG	120
20	CTCAGTGCGG CGGTGTGCCG GGCTGAGGCT GGGCTCGAAA CCGAAAGTCC CGTCCGGACC	180
	CTCCAAGTGG AGACCCTGGT GGAGCCCCCA GAACCATGTG CCGAGCCCGC TGCTTTTGGA	240
25	GACACGCTTC ACATACACTA CACGGGAAGC TTGGTAGATG GACGTATTAT TGACACCTCC	300
	CTGACCAGAG ACCCTCTGGT TATAGAACTT GGCCAAAAGC AGGTGATTCC AGGTCTGGAG	360
20	CAGAGTCTTC TCGACATGTG TGTGGGAGAG AAGCGAAGGG CAATCATTCC TTCTCACTTG	420
30	GCCTATGGAA AACGGGGATT TCCACCATCT GTCCCAGCGG ATGCAGTGCT GCAGTATGAC	480
	GTGGAGCTGA TTGCACTAAT CCGAGCCAAC TACTGGCTAA AGCTGGTGAA GGGCATTTTG	540
35	CCTCTGGTAG GGATGGCCAT GGTGCCAGCC CTCCTGGGCC TCATTGGGTA TCACCTATAC	600
	AGAAAGGCCA ATAGACCCAA AGTCTCCAAA AAGAAGCTCA AGGAAGAGAA ACGAAACAAG	660
40	AGCAAAAAGA AATAATAAATTTT AAAAAAAAA AAAAAAAA	717
45	(2) INFORMATION FOR SEQ ID NO: 27:(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 1099 base pairs	
	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
50	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 27:	
55	GGCACGAGCC GATGTGGACA TCATCCTGTC TATCCCCATG TTCCTGCGCC TGTACCTGAT	60
22	CGCCCGAGTC ATGCTGCTGC ACAGCAAGCT CTTCACCGAT GCCTCGTCCC GCAGCATCGG	120
	GGCCCTCAAC AAGATCAACT TCAACACCCG CTTTGTCATG AAGACGCTCA TGACCATCTG	180
60	CCCTGGCACT GTGCTGCTCG TGTTCAGCAT CTCTCTGTGG ATCATTGCTG CCTGGACCGT	240

	CCGTGTCTGT	GAAAGTCCTG	AATCACCAGC	CCAGCCTTCT	GGCTCATCAC	TTCCTGCTTG	300
5	GTACCATGAC	CAGCAGGACG	TAACTAGTAA	CTTTCTGGGT	GCCATGTGGC	TCATCTCCAT	360
3	CACATTCCTT	TCCATTGGTT	ATGGGGACAT	GGTGCCCCAC	ACATACTGTG	GGAAAGGTGT	420
	CTGTCTCCTC	ACTGGCATCA	TGGGTGCAGG	CTGCACTGCC	CTTGTGGTGG	CCGTGGTGGC	480
10	CCGAAAGCTG	GAACTCACCA	AAGCGGAGAA	GCACGTTCAT	AACTTCATGA	TGGACACTCA	540
	GCTCACCAAG	CGGATCAAGA	ATGCTGCAGC	CAATGTCCTT	CGGGAAACAT	GGTTAATCTA	600
15	TAAACACACA	AAGCTGCTAA	AGAAGATTGA	CCATGCCAAA	GTGAGGAAAC	ACCAGAGGAA	660
13	GTTCCTCCCA	AGCTATCCAC	CAGTTTGAGG	AGCGTCCCAG	ATGGAACAGA	GGAAAGCTGA	720
	GTGACCAAGC	CAACACTCTG	GTGGACCTTT	CCAAGATGCA	GAATGTCATG	TATGACTTAA	780
20	TCACAGAACT	CAATGACCGG	AGCGAAGACC	TGGAGAAGCA	GATTGGCAGC	CTGGAGTCGA	840
	AGCTGGAGCA	TCTCACCGCC	AGCTTCAACT	CCCTGCCGCT	GCTCATCGCC	GACACCCTGC	900
25	GCCAGCAGCA	GCAGCAGCTC	CTGTCTGCCA	TCATCGAGGC	CCGGGGTGTC	AGCGTGGCAG	960
23	TGGGCACCAC	CCACACCCCA	ATCTCCGATA	GCCCCATTGG	GGTCAGCTCC	ACCTCCTTCC	1020
	CGACCCCGTN	CACAAGTTCA	AGCAGTTGCT	AAATAAATCT	CCCCACTCCA	GAAGCATTAA	1080
30	ААААААААА	AAAAAAA					1099
35	(2) INFORMA	ATION FOR SE	EO TO NO 28	·			
	(2, 23, 3122			•			
	(i)	SEQUENCE CH (A) LEN	HARACTERIST: GTH: 941 ba				
40			E: nucleic				
40	(C) STRANDEDNESS: double (D) TOPOLOGY: linear						
	(xi)) SEQUENCE I	DESCRIPTION	: SEQ ID NO	: 28:		
45	NAME COOKS	1010001100	@1@@@@#==				
T J	AATTCGGCAG	AGAGCCAACC	GAGGGCGTTC	CTGTCGGGGC	TGCAGCGGCG	GGAGGGAGCC	60

CAGTGGAGGC GCCCTCCCGA AGCGCCACTG CCCATGCTGA CCACCCAGCC CTCCGGCTGC 120 TGATGTCATG AGTAACACCA CTGTGCCCAA TGCCCCCCAG GCCAACAGCG ACTCCATGGT 180 50 GGGCTATGTG TTGGGGCCCT TCTTCCTCAT CACCCTGGTC GGGGTGGTGG TGGCTGTGGT 240 AATGTATGTA CAGAAGAAAA AGCGGGTGGA CCGGCTGCGC CATCACCTGC TCCCCATGTA 300 55 CAGCTATGAC CCAGCTGAGG AACTGCATGA GGCTGAGCAG GAGCTGCTCT CTGACATGGG

AGACCCCAAG GTGGTACATG GCTGGCAGAG TGGCTACCAG CACAAGCGGA TGCCACTGCT 420 GGATGTCAAG ACGTGACCTG ACCCCCTTGC CCCACCCTTC AGAGCCTGGG GTYCTGGACT 480

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	GCCTGGGGCC CTGCCATCTG CTTCCCCTGC TGTCACCTGG STCCCCCTGC TGGGTGCTGG	540
	GTCTCCATTT CTCCCTCCAC CCACCCTCAG CAGCATCTGC TTCCCATGCC CTCACCATCA	600
5	CCTCACTGCC CCCAGGCCTT CTGCCCTTTG TGGGTGTTGA GCTCACCGCC CACCCACAGG	660
	CACTCATGGG AAGAGGCTTT CCTTCTGGGA TGGCGGCGGC TGGTAGACAC CTTTGCTTTC	720
10	TCTAGCCCTC CTGGGCTGGG CTTGGGCACA AATCCCCAGG CAGGCTTTGG AGTTGTTTCC	780
	ATGGTGATGG GGCCAGATGT ATAGTATTCA GTATATATTT TGTAAATAAA ATGTTTTGTG	840
	GCTAAAAAAA AAAAAAAAAA ATCNAAGGGG GGGCCGGTAC CCAAATTCCC CCTATANTGA	900
15	ATTCGTATTA ACAATTCACT TGGGGCCGTC CTTTTAANAA C	941
30	TO THE MENT TO THE TOTAL TO MO. 20.	
20	(2) INFORMATION FOR SEQ ID NO: 29:	
25	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 756 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 29:	
30	GGCACGAGGA AGCTGGAGCG GGCCGGCGGT GCAGTCACGG GGGAGCGAGG CCTGCTGGGC	60
	TTGGCAACGA GGGACTCGGC CTCGGAGGCG ACCCAGACCA CACAGACACT GGGTCAAGGA	120
25	GTAAGCAGAG GATAAACAAC TGGAAGGAGA GCAAGCACAA AGTCATCATG GCTTCAGCGT	180
35	CTGCTCGTGG AAACCAAGAT AAAGATGCCC ATTTTCCACC ACCAAGCAAG CAGAGCCTGT	240
	TGTTTTGTCC AAAATCAAAA CTGCACATCC ACAGAGCAGA GATCTCAAAG ATTATGCGAG	300
40	AATGTCAGGA AGAAAGTTTC TGGAAGAGAG CTCTGCCTTT TTCTCTTGTA AGCATGCTTG	360
	TCACCCAGGG ACTAGTCTAC CAAGGTTATT TGGCAGCTAA TTCTAGATTT GGATCATTGC	420
45	CCAAAGTTGC ACTTGCTGGT CTCTTGGGAT TTGGCCTTGG AAAGGTATCA TACATAGGAG	480
	TATGCCAGAG TAAATTCCAT TTTTTTGAAG ATCAGCTCCG TGGGGCTGGT TTTGGTCCAC	540
	AGCATAACAG GCACTGCCTC CTTACCTGTG AGGAATGCAA AATAAAGCAT GGATTAAGTG	600
50	AGAAGGGAGA CTCTCAGCCT TCAGCTTCCT AAATTCTGTG TCTGTGACTT TCGAAGTTTT	660
	TTAAACCTCT GAATTTGTAC ACATTTAAAA TTTCAAGTGT ACTTTAAAAT AAAATACTTC	720

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(2) INFORMATION FOR SEQ ID NO: 30:

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WO 98/42738

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186

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2100 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 30:

10	NCCAGAGGCA	GAAAGTCCTG	CTTCTGGGGC	GTAACCTACA	GGATATCCTT	GGAACAGAAG	60
10	ATCTTATTGT	GGAAGTRACT	TCCAATGATG	CTGTGAGATT	TTATCCCTGG	ACCATTGATA	120
	ATAAATACTA	TTCAGCAGAC	ATCAATCTAT	GTGTGGTGCC	AAACAAATTT	CTTGTTACTG	180
15	CAGAGATTGC	AGAATCTGTC	CAAGCATTTG	TGGTTTACTT	TGACAGCACA	CAAAAATCGG	240
	GCCTTGATAG	TGTCTCCTCA	TGGCTTCCAC	TGGCAAAAGC	ATGGTTACCY	GAGGTGATGA	300
20	TCTTGGTCTG	CGATAGAGTG	TCTGAAGATG	GTATAAACCG	ACAAAAAGCT	CAAGAATGGT	360
20	GCATCCAAAC	ATGGCTTTGA	ATTGGTAGAA	CTTAGTCCAG	AGGAGTTGCC	TGAGGAGGAT	420
	GATGACTTCC	CAGAATCTAC	AGGAGTAAAG	CGAATTGTCC	AAGCCCTGAA	TGCCAATGTG	480
25	TGGTCCAATG	TAGTGATGAA	GAATGATAGG	AACCAAGGCT	TTAGCTTGCT	GCAACTCATT	540
	GACTGGAACA	AACCATAGCA	TTGGGTCAGC	AGATCCCTGT	CACCCAGAGC	AACCCCATTT	600
30	GCCAGCAGCA	GATAGTACTG	AATCCCTCTC	TGATCATCGG	GGTGGTGCAT	CTAACACAAC	660
	AGATGCCCAG	GTTGATAGCA	TTGTGGATCC	CATGTTAGAT	CTGGATATTC	AAGAATTAGC	720
	CAGTCTTACC	ACTGGAGGAG	GAGATGTGGA	GAATTTTGAA	AGACTCTTTT	CAAAGTTAAA	780
35	GGAAATGAAA	GACAAGGCTG	CGACGCTTCC	TCATGAGCAA	AGAAAAGTGC	ATGCAGAAAA	840
	GGTGGCCAAA	GCATTCTGGA	TGGCAATCGG	GGGAGACAGA	GATGAAATTG	AAGGCCTTTC	900
40	ATCTGATGAA	GAGCACTGAA	TTATTCATAC	TAGGGTTTGA	CCAACAAAGA	TGCTAGCTGT	960
	CTCTGAGATA	CCTCTCTACT	CAGCCCAGTC	ATATTTTGCC	AAAATTGCCC	TTATCATGTT	1020
	GGCTGCCTGA	CTTGTTTATA	GGTCCCCTT	AATTTTAGTT	TTTAGTAGGA	GGTTAAGGAG	1080
45	AAATCTTTTT	TTTCCTCAGT	ATATTGTAAG	AGAGTGAGGA	ATACAGTGAT	AGTAATGAGT	1140
	GAGGATTTCT	TAAATRTACT	TTTTTTTTGT	TCTAGGAATG	AGGGTAGGAT	AAATCTCAGA	1200
50	GGTCTGTGTG	ATTTACTCAA	GTTGAAGACA	ACCTCCAGGC	CATTCCTGGT	CAACCTTTTA	1260
	AGTAGCATTT	CCAGCATTCA	CACTTGATAC	TGCACATCAG	GAGTTGTGTC	ACCTTTCCTG	1320
	GGTGATTTGG	GTTTTCTCCA	TTCAAGGAGC	TTGTAGCTCT	GAAGCTATGA	TGCTTTTATT	1380
55	GGGAGGAAAG	GAGGCAGCTG	CAGAATTGAT	GTGAGCTATG	TGGGGCCGAA	GTCTCAGCCC	1440
	GCAGCTAAGT	CTCTACCTAA	GAAAATGCCT	CTGGGCATTC	TTTTGAAGTA	TAGTGTCTGA	1500
60	GCTCATGCTA	GAAAGAATCA	AAAAGCCAGT	GTGGATTTTT	AGACTGTAAT	AAATGAGGCA	1560

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	AAGGATTTCT	ATTCCAGTGG	GAAGRAAACC	TCTCTACTGA	GTTGTGGGGG	ATATGTTGTA	1620
5	TGTTAGAGAG	AACCTTAAGG	AGTCCTTGTA	TGGGCCATGG	AGACAGTATG	TGATAACATA	1680
	CCGTGATTTT	CATGAAGAAA	TTCTTCTGTC	TTAGAGTTCT	CCCCTGCTGC	TTGAGATGCC	1740
	AGAGCTGTGT	TGTTGCACAC	CTGCAAAACA	AGGCACATTT	CCCCCTTTCT	CTTTAAAGCC	1800
10	AAAGAGAGAT	CACTGCCAAA	GTGGGAGCAC	TAAGGGGTGG	GTGGGGAAGT	GAAATGTTAG	1860
	GCGATGAATT	CCTGAGCACC	TIGITITICT	TCCAAGGTTC	GTAGCTCCTC	TCTGCCCTTC	1920
	CAAGCCTGTA	ACCTCGGAGG	ACTATCTTTT	GTTCTTTATC	CTTTGTCTTG	TTTGAGTGGG	1980
15	TCAGCCCCAG	AGGAACTGAT	AAGCAAATGG	CAAGTTTTTA	AAGGAAGAGT	GGAAAGTACT	2040
	GCAAATAAAA	ATCCTTATTT	GTTTTTGTAG	AAAAAAAAA	ААААААААА	AAAAAAAAAG	2100

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(2) INFORMATION FOR SEQ ID NO: 31:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1448 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

30 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 31:

	AAAAAAAAA	AAAGCCCACC	TGAAAGCCTG	TCTCTTTCCA	CTTTGTTGGC	CCTTCCAGTG	60
25	GGATTATCGA	GCATGTTGTT	TTTTCATAGT	GCCTTTTTCC	TTATTTCAAG	GGTTGCTTCT	120
35	GAGTGGTGTT	TTTTTTTTT	TTAATTTGTT	TTGTTTTAAA	ATAAGTTAAA	GACAGTCCAG	180
	AGCTTTTCAG	CCAATTTGTC	TCCTACTCTG	TGTAAATATT	TTTCCCTCCG	GGCAGGGGAG	240
40	CCAGGGTAGA	GCAAAGGAGA	CAAGCAGGAG	TGGAAGGTGA	GGCGTTCTCC	TGCTTGTACT	300
	AAGCCAGGAG	STTTAAGCTC	CAGCTTTAAG	GGTTGTGAGC	CCCTTGGGGT	TCAGGGAACT	360
45	GCTTGCCCAG	GGTGCAGTGT	GAGTGTGATG	GGCCACCGGG	GCAAGAGGGA	AGGTGACCGC	420
43	CCAGCTCTCC	CACATCCCAC	TGGATCTGGC	TTACAGGGGG	GTCGGAAGCC	TGTCCTCACC	480
	GTCTCGGGGG	TTGTGGCCCC	CGCCCCTCC	CTATATGCAC	CCCTGGAACC	AGCAAGTCCC	540
50	AGACAAGGAG	AGCGGAGGAG	GAAGTCATGG	GAACGCAGCC	TCCAGTTGTA	GCAGGTTTCA	600
	CTATTCCTAT	GCTGGGGTAC	ACAGTGAGAG	TACTCACTTT	TCACTTGTCT	TGCTCTTAGA	660
E E	TTGGGCCATG	GCTTTCATCC	TGTGTCCCCT	GACCTGTCCA	GGTGAGTGTG	AGGGCAGCAC	720
55	TGGGAAGCTG	GAGTGCTGCT	TGTGCCTCCC	TTCCCAGTGG	GCTGTGTTGA	CTGCTGCTCC	780
	CCACCCCTAC	: CGATGGTCCC	AGGAAGCAGG	GAGAGTTGGG	GAAGGCAAGA	TTGGAAAGAC	840
60	AGGAAGACCA	AGGCCTCGGC	: AGAACTCTCT	GTCTTCTCTC	CACTTCTGGT	CCCCTGTGGT	900

	GATGTGCCTG TAATCTTTTT CTCCACCCAA ACCCCTTCCC ACGACAAAAA CAAGACTGCC	960
5	TCCCTCTCTT CCGGGAGCTG GTGACAGCCT TGGGCCTTTC AGTCCCAAAG CGGCCGATGG	1020
	GAGTCTCCCT CCGACTCCAG ATATGAACAG GGCCCAGGCC TGGAGCGTTT GCTGTGCCAG	1080
	GAGGCGGCAG CTCTTCTGGG CAGAGCCTGT CCCCGCCTTC CCTCACTCTT CCTCATCCTG	1140
10	CTTCTCTTTT CCTCGCAGAT GATAAAAGGA ATCTGGCATT CTACACCTGG ACCATTTGAT	1200
	TGTTTTATTT TGGAATTGGT GTATATCATG AAGCCTTGCT GAACTAAGTT TTGTGTGTAT	1260
15	ATATTTAAAA AAAAAATCAG TGTTTAAATA AAGACCTATG TACTTAACTCTG	1320
13	CGGATAGCAT TTGGTAGGTA GTGATTAACT GTGAATAATA AATACACAAT GAATTCTTMA	1380
	AAAAAAAAAA AAAAAAAAAA AAAAAAAAAA AAACCCCGGG GGGGCCCCG GGCCCCAATT	1440
20	CCCCCCAA	1448
25	(2) INFORMATION FOR SEO ID NO: 32:	
23		
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 456 base pairs	
30	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 32:	
35	GGCACAGCAA ACTTGACGCC ATGAAGATCC CGGTCCTTCC TGCCGTGGTG CTCCTCTCCC	60
	TCCTGGTGCT CCACTCTGCC CAGGGAGCCA CCCTGGGTGG TCCTGAGGAA GAAAGCACCA	120
40	TTGAGAATTA TGCGTCACGA CCCGAGGCCT TTAACACCCC GTTCCTGAAC ATCGACAAAT	180
10	TGCGATCTGC GTTTAAGGCT GATGAGTTCC TGAACTGGCA CGCCCTCTTT GAGTCTATCA	240
	AAAGGAAACT TCCTTTCCTC AACTGGGATG CCTTTCCTAA GCTGAAAGGA CTGAGGAGCG	300
45	CAACTCCTGA TGCCCAGTGA CCATGACCTC CACTGGAAGA GGGGGCTAGC GTGAGCGCTG	360
	ATTCTCAACC TACCATAACT CTTTCCTGCC TCAGGAACTC CAATAAAACA TTTTCCATCC	420
50	AAAAAAAAA AAAAAAAAC CCCNGGGGGG GCCCGG	456
50		
	(2) INTEGRAMMITON FOR GEO TO NO. 22	
55	(2) INFORMATION FOR SEQ ID NO: 33:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1326 base pairs	
	(B) TYPE: nucleic acid	
60	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	

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	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 33:	
_	GGCACGAGTG CAGGCCCAGA GAGGACTCAT TGAAAGGACT GAAAGGGGAG GTGGCGTTTT	60
5	CTTCCTACCC AAACTTACCC CTGTGAGCTG GACAGCTTGG TAGCACCTGC CTGGACTTAG	120
	ATGGTGGTAG CCAAGAAGAC TGACATTTTA GGGAACAGGA CGGGGAGGAG AAGGCTCTGG	180
10	CACACACACA TGTGTCCATA TGTCCTGCAA TGGTCTGGGG ACTATTGCTA GGCTAGGAGC	240
	CCTAAGTGTC TTCTTCCTCA TGTCTMTTCT CCCCTGTSTC ATGGGCCCTA AGRTCTCTTT	300
15	CACTGGGCCT GCCTCAATGA ACGTGCTGCC CAGCTACCCC GAAACACGGC ANCTGCCGGC	360
13	TATCAATGCC CCAGCTGCAA TGGCCCATCT TCCCCCAACC AACCTGGCTG GGCCCGTGGG	420
	CTCCGCACTG AGARARAAS TTGGCACART CAACTGGGCC CGGGCAGGAC TGGGCCYCCC	480
20	TCTGATCGAT GAAGKTGGTG ARCCCAGAGC CCGAGCCCCT CAACACGTCT GACTTCTCTG	540
	ACTGGTCTAG TTTTAATGCC AGCAGTACCC CTGGACCAGA GGAGGTAGAC AGCGCCTCTG	600
25	CTGCCCCAGC CTTCTACAGC CGAGCCCCCC GGCCCCAGC TTCCCCAGGC CGGCCCGAGC	660
	AGCACACAGT GATCCACATG GGCAATCCTG AGCCCTTGAC TCACGCCCCT AGGAAGGTGT	720
	ATGATACGCG GGATGATGAC CGGACACCAG GCCTCCATGG AGACTGTGAC GATGACAAGT	780
30	ACCGACGTCG GCCGGCCTTG GGTTGGCTGG CCCGGCTGCT AAGGAGCCGG GCTGGGTCTC	840
	GGAAGCGRCC GCTGACCCTG CTCCAGCGGG CGGGGCTGCT GCTACTCTTG GGACTGCTGG	900
35	GCTTCCTGGC CCTCCTTGCC CTCATGTCTC GCCTAGGCCG GGCCGCAGCT GACAGCGATC	960
	CCAACCTGGA CCCACTCATG AACCCTCACA TCCGCGTGGG CCCCTCCTGA GCCCCCTTGC	1020
	TTGTGGCTAG GCCAGCCTAG GATGTGGGTT CTGTGGAGGA GAGGCGGGGT AATGGGGAGG	1080
40	CTGAGGGCAC CTCTTCACTG CCCCTCTCCC TCAAGCCTAA GACACTAAGA CCCCAGACCC	1140
	AAAGCCAAGT CCACCAGAGT GGCTGCAGGC CAGGCCTGGA GTCCCCGTGG GTCAAGCATT	1200
45	TGTCTTGACT TGCTTTCCTC CCGGGTYTCC AGCCTCCGAC CCCTCGCCCC ATGAAGGAGC	1260
	TGGCAGGTGG AAATAAACAA CAACTTTATT AAAAAAAAAA	1320
	AAANAA	1340

50

(2) INFORMATION FOR SEQ ID NO: 34:

55 (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 710 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 34:	
	GCGAAAGAGA AAAAGGCTGG AGCTCCCGCC CCCGGGGCTG TCAGATGGCT TGGGTTTCTG	60
5	CGACGCGATT GGCTCGCGGA GGGCAGAAAT TACTCAGCAA ACATGACTAT TATTAGCTGC	120
	TTAGCAACAG CTCACCAAAG TAGAGAGACC ACCCAGGTAG GCAACCCAGT GTGTGCATCC	180
10	TCGGCTTCGG GGCAGCCTCT GAGAGCGCCA ACCTTCTCGC ATGCAATACT TCCATTAAGG	240
10	AATGCTCCCC CTCCTTTCTC TCTTATTCCT TTTCTTTTCA ACAGTGTCTT CTTTTTGTGG	300
	GATGCCTTTG CGCGCACACA CGCGCGCGCA SGCACACACA CGAACATTTG CCTCGCGGTA	360
15	GACACGGGG GAAATGTWAT ATTTTTTTAA GCGCTTAAAC AATTTCTGAA ATTCCTCAAA	420
	GAAAAGCCTT TCAGARGCAC CTTGGCCTCA AGCTGCAACA AATACTGGGA RGTCCGGCTC	480
20	GCATTCCCAG GCCTGCACCA ATAATGACAG CGTGCTGGAT ARTGCGCCAG TGTGTGCCAG	540
20	ATTITITIT CCTCTTCTCT TTTCTTTTAT AACTAAAGGG AAGACTTAGG CTCTTGCAGG	600
	GAACAACGCC TCGCATTAAG ATAAACAGAA TGGAAAGTTA AAGAGGAAAG CAAGGACGTT	660
25	GGGAAAAGCC ATCTTTCTTA AAATCCGTCT GCCCCCCAGC CGCTTTCTCC	710
30	(2) INFORMATION FOR SEQ ID NO: 35:	
	(i) CEOLEMAN CHENTER CONTROL	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1188 base pairs	
35	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 35:	
40	GATGGCTTTT ATATCTATTA TCGACCCACA GACAGTGACA ATGATAGTGA CTACAAGAAG	60
	GATATGGTGG AAGGGGACAA GTACTGGCAC TCCATCAGCC ACCTGCAGCC AGAGACCTCC	120
45	TACGACATTA AGATGCAGTG CTTCAATGAA GGAGGGGAGA GCGAGTTCAG CAACGTGATG	180
	ATCTGTGAGA CCAAAGCTCG GAAGTCTTCT GGCCAGCCTG GTCGACTGCC ACCCCCAACT	240
	CTGGCCCCAC CACAGCCGCC CCTTCCTGAA ACCATAGAGC GGCCGGTGGG CACTGGGGCC	300
50	ATGGTGGCTC GCTCCAGCGA CCTGCCCTAT CTGATTGTCG GGGTCGTCCT GGGCTCCATC	360
	GTTCTCATCA TCGTCACCTT CATCCCCTTC TGCTTGTGGA GGGCCTGGTC TAAGCAAAAA	420
55	CATACAACAG ACCTGGGTTT TCCTCGAAGT GCCCTTCCAC CCTCCTGCCC GTATACTATG	480
55	GTGCCATTGG GAGGACTCCC AGGCCACCAG GCAGTGGACA GCCCTACCTC AGTGGCATCA	540
	GTGGACGGC CTGTGCTAAT GGGATCCACA TGAATAGGGG CTGCCCCTCG GCTGCAGTGG	600
	Tarming of Crockers	

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	CCAGCAGCCT	GCTGAGGCAG	ACCCATCTTG	GCAATGGATA	TGACCCCCAA	AGTCACCAGA	720
~	TCACGAGGGG	TCCCAAGTCT	AGCCCGGACG	AGGGCTCTTT	CTTATACACA	CTGCCCGACG	780
3	ACTCCACTCA	CCAGCTGCTG	CAGCCCCATC	ACGACTGCTG	CCAACGCCAG	GAGCAGCCTG	840
	CTGSTGTGGG	CCAGTCAGGG	GTGAGGAGAG	CCCCGACAG	TCCTGTCCTG	GAAGCAGTGT	900
10	GGGACCCTCC	ATTTCACTCA	GGGCCCCCAT	GCTGCTTGGG	CCTTGTGCCA	GTTGAAGAGG	960
	TGGACAGTCC	TGACTCCTGC	CAAGTGAGTG	GAGGAGACTG	GTGTCCCCAG	CACCCCGTAG	1020
15	GGGCCTACGT	AGGACAGGAA	CCTGGAATGC	AGCTCTCCCC	GGGCCACTG	GTGCGTGTGT	1080
13	CTTTTGAAAC	ACCACCTCTC	ACAATTTAGG	CAGAAGCTGA	TATCCCAGAA	AGACTATATA	1140
	TTGTTTTTT	TTTAAAAAAA	AAAAAAAA	AWCYCGGGGG	GGGGCCC		1188
20							

20

(2) INFORMATION FOR SEQ ID NO: 36:

(i) SEQUENCE CHARACTERISTICS: 25

- (A) LENGTH: 956 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: double
- (D) TOPOLOGY: linear

30

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36:

	GGCAGAGCAG	TGAAAATGCA	TCCTAAAAAT	TCAATGTTTA	TACCAGGCTC	ATGACACTAA	60
35	GATGTGACAT	CTGGACACGA	GGGGTCAGCC	ACGTGGATAC	ATCCCTCCCA	GATTGCATCT	120
	CCAGGAATCA	CTCTGCTAGC	AGAATGGGCG	CCCCATCCCT	TACTATGCTG	CTCCTCCTCA	180
40	AAGTGCAGCC	CAGAAGGACC	CAGGCCTTTG	ATGCACATTG	GGTGGGTCTC	CCACTACTTT	240
40	AGTTGAAATG	GGAGCATGCT	GGAGTCGGCG	TTCTGTTGCT	TCTGGTGAGA	AGGACATCCC	300
	ATTGACCCCT	GGCCACCAGG	TCCAGTATTC	CATCCTTCCT	TCTGTCCCAG	CCTATCGCCC	360
45	TCCCCACYAG	GCCCACCCC	ACAACTTCTC	CTCAAGGGAG	GTTNTCCCGC	AGCTGGAGGG	420
	CTTGCACAGA	CCAGCAGTCA	CAGAAATCAT	TCTTCCTGCT	GTACTGGGCC	TTAACTGCCT	480
50	GCAAATGTCC	GAGCACTACT	GCATAGGATG	CCAGAGCCAC	CGAAGATAAA	CACAGCCAAG	540
50	ATAATAATTT	ATAAAAGGAA	AAATCTCAGC	CTGCAGAACT	CTGGTTTTGA	CCCACCATCG	600
	GCCAGATGCA	CATCTTCAGG	GCCTGTTGAG	CACCTTCTGA	AAAGCAGGGC	TCGTAATAGA	660
55	CTCCAGCACA	TTCCATCAGA	. GTCAGGAAAA	CTGCGGTGAG	TCCCAGAGAA	TCTAGGGTGC	720
	AGGGCAGGGA	GCAGGAGTCA	TAAGGAGTGA	TAACCTAAAC	TGTGTGTAGT	CAGCGGGGAG	780
60	GGTCTTATGT	TATCAGGTGA	AATGAGAGCC	: AGTAAGTTAG	TTGATCCTGT	CACAGATATA	840

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	ACCCIGATAA	CACCCCATAG	ATACGCGACA	CGTGTGTCCT	GCCCCTGCTT	TCCCCATCCA	900
	ACATGGTTCT	TCTGTTCCAC	AGACATTAAA	GGGCTTTCT	GCAATTACTT	АААААА	956
5							
	(2) TITTOPI			_			
10		ATION FOR S					
10	(i)	(B) TYP (C) STR	HARACTERIST GTH: 1603 b E: nucleic ANDEDNESS: OLOGY: line	ase pairs acid double			
	(xi) SEQUENCE	DESCRIPTION	: SEQ ID NO	: 37:		
	TCGACCCACG	CGTCCGCTCT	GCCAGGAATC	TGGTCTTTCT	GTAGACCCAA	GTCAGAAAGA	60
20	ACCATTTGTG	GAGTTAAATC	GAATATTAGA	RGCATTAAAR	GTCAGAGTTC	TGAGACCTGC	120
	TCTGGAATGG	GCAGTTICAA	ACCGAGAGAT	GCTTATAGCC	CAAAACAGCT	CCTTGGAATT	180
25	TAAACTACAC	AGACTGTATT	TTATTAGCTT	RTTAATGGGT	GGAACACAAA	TCAGCGAGAR	240
23	GCATTACAAT	ATGCTAAAAA	TTTTCAGCCA	TTTGCCCTAA	ATCATCAAAA	AGACATTCAG	300
	GTTTTGATGG	GAAGCCTTGT	GTACCTGAGA	CAAGGGATTG	AGAACTCACC	ATATGTTCAC	360
30	CTACTTGATG	CAAACCAGTG	GGCTGATATC	TGTGACATCT	TTACACGGGA	TGCTTGTGCC	420
	CTCCTGGGGC	TCTCCGTGGA	GTCCCCTCTC	AGTGTCAGTT	TCTCAGCAGG	TTGTGTGGCG	480
35	CTGCCAGCTT	TAATTAACAT	CAAAGCCGTG	ATTGAACAGA	GGCAGTGTAC	TGGAGTTTGG	540
33	AACCAGAAAG	ATGAATTACC	TATTGAAGTG	GACCTTGGTA	AAAAGTGCTG	GTATCACTCT	6 0 0
	ATATTTGCCT	GCCCCATTCT	TCGTCAGCAA	ACAACAGATA	ACAATCCACC	CATGAAATTG	660
40	GTCTGTGGTC	ATATTATATC	AAGAGATGCC	CTGAATAAAA	TGTTTAATGG	TAGCAAATTA	720
	AAATGTCCCT	ACTGTCCAAT	GGAACAAAGT	CCAGGAGATG	CCAAACAGAT	ATTTTTCTGA	780
15	AGAGATAACT	TTAGTTTGCA	ATTTGTAAGT	GAAACTGAAT	CGTGGGTGCA	TTTCAGAAGA	840
45	GAACGTTCCA	TATAATGCAG	CTAACCAAGG	ACTCCTGTGT	TTCTATAAGC	TAATGCTCCA	900
	GAAACTTTGC	CAACCTGTTA	GTGTACACAC	ACTGAGGGGA	GTGCTCCCGG	TGAATATTAT	960
50	CATAGGGCTT	TATTATATTC	TTGGTCTTCA	TTTCTGATCA	AGTAAATACA	CCAGCAGTTG	1020
	TCATTCAATG	CAGGTTTTTG	TACTTAATTA	TATGGTGATT	TTTTTACTTT	TTAAGAGCAG	1080
	AAACGGAAAT	TGACCTCCCC	GCCATGTGTT	TAATATTCCT	CCTGCTTTTA	CTTTTGTCAT	1140
55				TTGTGAAAAA			1200
						GGTTATGAAA	1260
60				AAAGCTGTGA			1320

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	TGATGTTTTC	CTCTGCTCCA	GCTCCAAGAA	GTCAGCACAC	CTGCATTTTA	GCTCTGCATG	1380
5	CAGCCCCAGC	AGGCTGCGTG	TTTAAGAATT	TCATTGTTTA	ACTGGCTGGT	GTGAGAAGTC	1440
	TTCCGTTAGC	ATAGAGTGGA	AGGAGTACTA	TTGTTTGGTT	GGGTTTTTGT	TTGTTTGTTT	1500
	TTTGTTTTTG	CTTTTATTGC	CAAGAGGTGC	TTGTTTTAAA	AGTATGTTTA	ATAAAATGAA	1560
10	ATTCTAAAGT	TAARAAGTGT	TCTTAAAGTT	GATATTTAAC	TCT		1603

(2) INFORMATION FOR SEQ ID NO: 38:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1089 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 38:

GGCACGAGCT ACCTTTCTGC CTGCTTTGCT GGCTGCAACA GCACGAATCT CACGGGCTGT 60 25 GCGTGCCTCA CCACCGTCCC TGCTGAGAAC GCAACCGTGG TTCCTGGAAA ATGCCCCAGT 120 CCTGGGTGCC AAGAGGCCTT CCTCACTTTC CTCTGTGTGA TGTGTATCTG CAGCCTGATC 180 30 GGTGCCATGG CAAGACACCC TCAGTCATCA TCCTCATCAG GACAGTCAGC CCTGAACTCA 240 AGTOTTACGO TTTGGGAGTT CTTTTTCTCC TCCTTCGTTT GTTGGGCTTC ATCCCTCCAC 300 CCCTCATCTT CGGGGCTGGC ATCGACTCCA CCTGCCTGTT CTGGAGCACG TTCTGTGGGG 360 35 AGCAAGGCGC CTGCGTCCTC TACGACAATG TGGTCTACCG ATACCTGTAT GTCAGCATCG 420 480 CCATCGCGCT CAAATCCTTC GCCTTCATCC TGTACACCAC CACGTGGCAG TGCTGAGGAA 40 AAACTATAAA CGCTACATCA AAAACCACGA GGGCGGGCTG AGCACCAGTG AGTTCTTTGC 540 CTCTACTCTG ACCCTAGACA ACCTGGGGAG GGACCCTGTG CCCGCAAACC AGACACATAG 600 GACAAAGTTT ATCTATAACC TGGAAGACCA TGAGTGGTGT GAAAACATGG AGTCCGTTTT 45 660 720 ATAGTGACTA AAGGAGGGCT GAACTCTGTA TTAGTAATCC AAGGGTCATT TTTTTCTTAA AAAAAGAAAA AAAGGTTCCA AAAAAAACCA AAACTCAGTA CACACACACA GGCACAGATG 50 CACACACAC CAGACAGACA CACCGACTTT GTCCTTTTTC TCAGCATCAG AGCCAGACAG 840 GATTCAGAAT AAGGAGAGAA TGACATCGTG CGGCAGGGTC CTGGAGGCCA CTCGCGCGGC 900 TGGGCCACAG AGTCTACTTT GAAGGCACCT CATGGTTTTC AGGATGCTGA CAGCTGCAAG 960 55 CAACAGGCAC TGCCAAATTC AGGGAACAGT GGTGGCCAGC TTGGAGGATG GACATTTCTG 1020 1080 GATACACATA CACATACAAA ACAGAAAACA TTTTTTAAAA GAAGTTTCCT AAAATAAAAA 60

	AAAAAAAA	108
5	(2) INFORMATION FOR SEQ ID NO: 39:	
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 629 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 39:	
	AGCTCAGTTC CCTTAGAAAT GAAATTTTAA ATGACACTAC CAGGTAAGCC ACTGAGACCA	60
	GTGGAGGTGA TAGCTAAGAA CATAAGGAAT TAAGAATTTT TAATGGAGAA AGGAGGTAAT	120
20	GAATACCAGT TACATCCTAA GACTCACTGT AGTGGTGAGT GTTGTAATTT ATCTCGCTAT	180
	CCATCCTCTT TTAAGTTTTT CCTTAGAAAG TCCTCTATTG GTACCTTGGA GGGACTGCTG	240
25	TCAAAATATA TGGAAAAGTG GGTCTGTGTG GTACAAGAGG TGGACTTTGC CACACATGGA	300
23	AGTTTGCTGC CAAGATCTTC ACTAATGAAA GAAATCACCA GTGAGCTGCA CAGATTAGCC	360
	AAATACTGAG CTCATTAGAA CTACTAAGGC CTGGACATTT CTGCCTAATC CAGGACTCCT	420
30	GTAATTATCA GTCTTTGCTT TGGAGCTTCC CATTGTGTAG CTGARAATTT GTCATATCTG	480
	CATTATAATC TAAGGCTCCA CATACTTAAT CCTGCTTCTC CCCCTTTTTC TTTCCCTTTC	540
	CCAGCGGTCA GCTCTGCTGC ATAGTCTGAA GACTTTCCCT GCCCAATCCT GATAAAATTC	600
35	TTGCACTCGT AACCCCATCT CAGTGTCTG	629
40	(2) INFORMATION FOR SEQ ID NO: 40:	02 .
45	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1964 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 40:	
50	AAGAAGACAT GGAAATTGCT GAAGGATGTT TCAGGCATAT TAAGAAAATC TTTACGCAGC	60
	TTGAGGAATT CAGAGCCTCT GAATTGCTTC GAAGTGGACT GGACAGATCT AAATACCTTT	120
55	TAGTGAAAGA AGCCAAAATT ATTGCTATGA CCTGTACTCA TGCTGCCTTA AAACGACATG	180
	ACTTGGTCAA GCTAGGTTTC AAGTATGACA ACATTTTGAT GGAAGAGGCT GCTCAGATTC	240
.	TGGAGATAGA AACTITTATC CCTCTTCTTC TACAGAATCC TCAGGATGGA TTTAGCCGAC	300
60		

	TAAAACGATG GATTATGATT GGCGATCATC ACCAGTTACC TCCAGTTATT AANGAACATG	360
	GCCTTTCAAA AGTACTCAAA CATGGAGCAG TCTCTCTTCA CTCGCTTTGT TCGCGTTGGA	420
5	GTTCCGACTG TTGACCTTGA TGCTCAAGGG AGAGCCAGAG CAAGCTTGTG CAMCTNCTAC	480
	AACTGGCGAT ACAAGAATCT AGGAAACTTA CCCCATGTGC AGCTCTTGCC AGAGTTTAGT	540
4.0	ACAGCAAATG CTGGCTTACT GTATGACTTC CAGCTCATTA ATGTTGAAGA TTTTCAAGGA	600
10	GTGGGAGAAT CTGAACCTAA TCCTTACTTC TATCAGAATC TTGGAGAGGC AGAATATGTA	660
	GTAGCACTTT TTATGTACAT GTGTTTACTT GGTTACCCTG CTGACAAAAT CAGTATTCTA	720
15	ACAACATATA ATGGCCAAAA GCATCTTATT CGCGACATCA TCAATAGACG ATGTGGAAAC	780
	AATCCATTGA TTGGAAGACC AAACAAGGTG ACAACTGTTG ATAGATTTCA AGGTCAACAG	840
20	AATGACTATA TTCTTCTTTC TCTGGTACGA ACCAGGGCAG TGGGCCATCT GAGGGATGTC	900
20	CGTCGCTTGG TAGTGGCCAT GTCTAGAGCC AGACTTGGAC TTTATATCTT CGCCAGAGTA	960
	TCCCTCTTCC AAAACTGTTT TGAACTGACT CCAGCTTTCA GTCAGCTCAC AGCTCGCCCC	1020
25	CTTCATTTGC ATATAATTCC AACAGAACCT TTCCCAACTA CTAGAAAGAA TGGAGAGAGA	1080
	CCATCTCATG AAGTACAAAT AATAAAAAAT ATGCCCCAGA TGGCAAACTT TGTATACAAC	1140
20	ATGTACATGC ATTTGATACA GACTACACAT CATTATCATC AGACTTTATT ACAACTACCA	1200
30	CCTGCTATGG TAGAAGAGGG TGAGGAAGTT CAAAATCAAG AAACAGAATT GGAAACAGAA	1260
	GAAGAGGCCA TGACTGTTCA AGCTGACATC ATACCCAGTC CAACAGACAC CAGCTGCCGT	1320
35	CAAGAAACTC CAGCCTTTCA AACTGACACC ACCCCCAGTG AGACAGGAGC CACTTCCACT	1380
	CCAGAAGCCA TCCCTGCTTT ATCTGAGACC ACCCCTACTG TGGTAGGAGC TGTATCTGCA	1440
40	CCGGCAGAAG CTAACACACC TCAGGATGCC ACATCTGCCC CAGAAGAGAC CAAGTAGCCA	1500
40	AACTGTAGTC CTTCTAAAGG AGGACATGGC AGTCAAAAAG TCTGAGTAAA GCTGTTTTTT	1560
	GTATTTTATA TITGCTTCTG CCATTTTACT GTCACTAATT AATGTTTAGT TCTTATATTT	1620
45	GTTAACTGAT TTCGGTGTCT TGAATATATT TTTTTAAATT ATGTGTATGA ACAATTCTAG	1680
	TTTCATTTGT TCAATCAGAA GAGCAAATAA CCATTCCTTT CATGTTTTGA TCACTGAGTG	1740
50	TGTCTGTAAT CATACCTACA TTAAAATCAT TTTCTATGAA TATATAATAT ATACTTCACA	1800
50	TTTTTAGTGA ACTTCTCTAA AGAAGAGGAC AGAATATACT GGACTTAACC ACGAATACCC	1860
	TTGAGTGTCC AAATTGGGAA GGAACTKGTT TCTTCYGTTA TACTAYCAAA TGCTTAAATT	1920
55	CKGTTTCCTT TTTTCTTACC TTTGTTTGCT GTCTTTATGT AAAG	196

60 (2) INFORMATION FOR SEQ ID NO: 41:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1522 base pairs

(B) TYPE: nucleic acid(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 41:

	(31.2	, properties :	DESCRIPTION	. SEQ ID NO	. 41.		
10	CGTGTCCGCG	CGCCTGGGAG	ACGCTGCCTC	GGCCCGGACG	ceccecec	CCCGCGGCTG	60
	GAGGGTGGTC	GCCACTGGGA	CACTGTGAAC	CAGGAGTRAG	TCGGAGCTGC	CGCGCTGCCC	120
15	AGGCCATGGA	CTGTGAGGTC	AACAACGGTT	CCAGCCTCAG	GGATGAGTGC	ATCACAAACC	180
15	TACTGGTGTT	TGGCTTCCTC	CAAAGCTGTT	CTGACAACAG	CTTCCGCAGA	GAGCTGGACG	240
	CACTGGGCCA	CGAGCTGCCA	GTGCTGGCTC	CCCAGTGGGA	GGGCTACGAT	GAGCTGCAGA	300
20	CTGATGGCAA	CCGCAGCAGC	CACTCCCGCT	TGGGAAGAAT	AGAGGCAGAT	TCTGAAAGTC	360
	AAGAAGACAT	CATCCGGAAT	ATTGCCAGGC	ACCTCGCCCA	GGTCGGGGAC	AGCATGGACC	420
25	GTAGCATCCC	TCCGGGCCTG	GTGAACGGCC	TGGCCCTGCA	GCTCAGGAAC	ACCAGCCGGT	480
20	CGGAGGAGGA	CCGGAACAGG	GACCTGGCCA	CTGCCCTGGA	GCAGCTGCTG	CAGGCCTACC	540
	CTAGAGACAT	GGAGAAGGAG	AAGACCATGC	TGGTGCTGGC	CCTGCTGCTG	GCCAAGAAGG	600
30	TGGCCAGTCA	CACGCCGTCC	TTGCTCCGTG	ATGTCTTTCA	CACAACAGTG	AATTTTATTA	660
	ACCAGAACCT	ACGCACCTAC	GTGAGGAGCT	TAGCCAGAAA	TGGGATGGAC	TGAACGGACA	720
35	GTTCCAGAAG	TGTGACTGGC	TAAAGCTCGA	TGTGGTCACA	GCTGTATAGC	TGCTTCCAGT	780
55	GTAGACGGAG	CCCTGGCATG	TCAACAGCGT	TCCTAGAGAA	GACAGGCTGG	AAGATAGCTG	840
	TGACTTCTAT	TTTAAAGACA	ATGTTAAACT	TATAACCCAC	TTTAAAATAT	CTACATTAAT	900
40	ATACTTGAAT	GAAAATGTCC	ATTTACACGT	ATTTGAATGG	CCTTCATATC	ATCCACACAT	960
	GAATCTGCAC	ATCTGTAAAT	CTACACACGG	TGCCTTTATT	TCCACTGTGC	AGGTTCCCAC	1020
45	TTAAAAATTA	AATTGGAAAG	CAGGTTTCAA	GGAAGTAGAA	ACAAAATACA	ATTTTTTTGG	1080
,,,	TAAAAAAAA	TTACTGTTTA	TTAAAGTACA	ACCATAGAGG	ATGGTCTTAC	AGCAGGCAGT	1140
	ATCCTGTTTG	AGGAAAGCAA	GAATCAGAGA	AGGAACATAC	CCCTTACAAA	TGAAAAATTC	1200
50	CACTCAAAAT	AGGGACTATC	YATCTTAATA	CTAAGGAACC	AACAATCTTC	CTGTTTAAAA	1260
	AACCACATGG	CACAGAGATT	CNGAACTAAA	GTGCTGCACT	CAAATGATGG	GAAGTCCCGG	1320
55	CCCCAGTACA	CCAGGGGCTT	TGGACTTTTT	TCAACTTCGT	TTCCTTTTGT	TTGGANTCCA	1380
33	AAAGAACCAC	TTTGTGGTTC	TTAAAAGGGT	GTGAAGGTGA	TTTAAGGGC	CCAGGTCAGC	1440
	CACTGGTTGG	TTTACAAAAT	CNGGGTAACT	AACTGCATAC	AACTTTTTCC	CNTTTCCATG	1500
60	NCATCAGGAC	TTTGCTAAAG	AC				1522

5	(2) INFORMATION FOR SEQ ID NO: 42:	
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 875 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 42:	
15	TGGGATTTCC CTTTATCATG GAGGCCTTGT CCCACTTCCT CTATGTCCCT TTCCTTGGTG	60
	TCTGTGTCTG TGGGGCCATC TACACTGGCC TGTTCCTTCC TGAGACCAAA GGCAAGACCT	120
20	TCCAAGAGAT CTCCGAGGAA TTACACAGAC TCAACTTCCC CAGGCGGGCC CAGGGCCCCA	180
20	CGTGGAGGAG CCTGGAGGTT ATCCAGTCAA CAGAACTCTA GTCCCAAAGG GGTGGCCGTA	240
	GCCAAAGCCA GCTACCGTCC TGTCCTCTGC TTCCTGCCAG GGCCCTGGTC CTCAMTYCCT	300
25	YCTGCATTCC TCATTTAAGG AGTGTTTATT GAGCACCCTT TGTGTGCAGA CATGGCTCCA	360
	GGTGCTTAGC AATCAWTGGT GAGCGTGGTA TCCAGGCTAA AGGTAATTAA CTGACAGRAA	420
20	ATCAGTAACA ACATAATTAC AGGYTGGTTG TGGCAGYTCA TGACTGTAAT CCCAGCACTT	480
30	TTGGGAGCCA AGGTGGGARG ATCAATTGAG GCCAGAGTTT GAAAMCAGCT AGGTAACATA	540
	GTGAGACCCC CTATCTCTAC AAAAAATTTT AAACATTAGC TGGGCATGGT GGTATGTGCT	600
35	AACAGCTCTA GCTACTCAGG AGGCTGAGGC AGCAGGATCA CTTGAGTCCA AGAGTTCAAG	660
	GTAGCAGTAA GCTACAATCA CACCACTGCA TGCCAGACTG GGTGACAGAG GGAGACTTCA	720
40	TCTCTTTAAA ACATAATAAT AATAATTACA GACTCAGGAA ATGCAGTGAA AGAAAAATAC	780
40	AGGTTGGCCA GGTGAGGTGG CTGATGCCTG TAATCCCAGC ACTTTGGGAG GCCAAGATGG	840
	GAAGATTGCT TTGAGACCAG AAGTTTGAGA CCAGC	875
45		
	(2) INFORMATION FOR SEQ ID NO: 43:	
50 55	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 843 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
23	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 43:	
	CCCACGCGGT CCGNATCGTC CTTCCCTCAC TTCAGAGGGT GGCCAGAGCT GAATACCCAG	60
60	AGAGGGACAA GTAAGGGTCC AGTTCCAAAA CATCATGAGG ATGTATCATC CCACGTGTCT	120

	CACCTGACAG T	TTACAGAGGA	AACCCGCACC	CAGAATGCAC	GTGCTGTCTT	ATGGGAACAC	180
5	TCAGCGCAGA C	GTGCTCAGGT	CCGGCCACAC	TCGGGCTGTG	CTTGGTCGTG	CCATGGAATT	240
J	CCTCAGGACT T	TCTCAGCCT	CCCTAATGGC	AGAAGCCCCT	TTACAGCAAG	ACATTTACCG	300
	TTTGTCTGAA A	AATAGCCGAA	CTGAGCTTTT	CTTCAGGCTA	TATGAGAAGT	CTCTAGACAG	360
10	TGGGCACCGT C	CAGAAAGCCC	AGAGCCTTGT	GATAGCTCCC	ACCCTGCCTG	GCTCAGATCT	420
	TCCCATTTTT 1	TTTCCTCTGG	CACTAACCTC	ACCTTTTGTT	TTTTTGTGTT	TGTGTTTGTT	480
15	TTTGTTTTTG C	CAGAGTTGGA	TTACAGAAAC	TCCTATGAAA	TTGAATATAT	GGAGAAAATT	540
10	GGCTCCTCCT T	PACCTGTAAG	TTCGTCTGCC	TCGGGCCACT	TAGGGGACTC	GCTTTCCTGC	600
	CTTCAGGGGC C	CTCCTCCCCT	GTGCAGAGTG	TCTCTGGGAG	CTCAGACCCC	AAATCGAGTG	660
20	TTTTCTGTGT A	ACACAGCTTC	CCGGGTGCAC	AGCAATGATG	GACTGGGGCT	GGGGGTTGA	720
	GGTTTGTACT C	CAATCCACTT	CG TTT GACAT	TTTCAGGGAG	AAAATGATAG	AATACAATTA	780
25	GACGTCCTGC A	AGAATTACTT	TCCTAGACTG	AGAAAGAGCT	AGAGATTTCT	TTAAAAAAA	840
	AAA						843
30	(2) INFORMAT	TION FOR SE	Q ID NO: 44	i:			
35	(i) S	(A) LENG (B) TYPI (C) STRA	ARACTERIST: GTH: 489 base: E: nucleic and ANDEDNESS: O DLOGY: line	se pairs acid double			
40	(xi)	SEQUENCE I	DESCRIPTION	SEQ ID NO	: 44:		
+0	CTCTTAGGCT T	TTGAAGCATT	TTTGTCTGTG	CTCCCTGATC	TTCAGGTCAC	CACCATGAAG	60
	TTCTTAGCAG T	CCTGGTACT	CTTGGGAGTT	TCCATCTTTC	TGGTCTCTGC	CCAGAATCCG	120
45	ACAACAGCTG C	CTCCAGCTGA	CACGTATCCA	GCTACTGGTC	CTGCTGATGA	TGAAGCCCCT	180
	GATGCTGAAA C	CACTGCTGC	TGCAACCACT	GCGACCACTG	CTGCTCCTAC	CACTGCAACC	240
50	ACCGCTGCTT C	CTACCACTGC	TCGTAAAGAC	ATTCCAGTTT	TACCCAAATG	GGTTGGGGAT	300
30	CTCCCGAATG G	STAGAGTGTG	TCCCTGAGAT	GGAATCAGCT	TGAGTCTTCT	GCAATTGGTC	360
	ACAACTATTC A	ATGCTTCCTG	TGATTTCATC	CAACTACTTA	CCTTGCCTAC	GATATCCCCT	420
55	ттатстстаа т	CAGTTTATT	TTCTTTCAAA	TAAAAAATAA	CTATGAGCAA	САААААААА	480

199

	(2) INFORMATION FOR SEQ ID NO: 45:	
5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 534 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 45:	
	GAAGCAGTGT GTATCTATGA TTATATCTCT GTTCATCTAT ATATTTTTGA CATGTAGCAA	60
1.5	CACCTCTCCA TCTTATCAAG GAACTCAACT CGGTCTGGGT CTCCCCAGTG CCCAGTGGTG	120
15	GCCTTTGACA GGTAGGAGGA TGCAGTGCTG CAGGCTATTT TGTTTTTTGT TACAAAACTG	180
	TCTTTTCCCT TTTCCCCTCC ACCTGATTCA GCATGATCCC TGTGAGCTGG TTCTCACAAT	240
20	CTCCTGGGAC TGGGCTGAGG CAGGGGCTTC GCTCTATTCT CCCTAACCAT ACTGTCTTCC	300
	TTTCCCCTTG CCACTTAGCA GTTATCCCCC CAGCTATGCC TTCTCCCTCC CTCCCTTGCC	360
25	CTGGCATATA TTGTGCCTTA TTTATGCTGC AAATATAACA TTAAACTATC AAGTGAAAAA	420
23	AAAAAAAAA AAAACTCCAA GGGGGGGCCG GTACCCAATT CCCCCTATAN TGAGTCNTAT	480
	TACAATTCAC TGGGCCGTCG TTTTACAACG TCGTGAATGG GAAAACCTGG GCGT	534
30		
	(2) INFORMATION FOR SEQ ID NO: 46:	
35	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1374 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 46:	
	GGCACGAGTC CGGGATGAGC TCAGCCGCGG CCGACCACTG GGCGTGGTTG CTGGTGCTCA	60
45	GCTTCGTGTT TGGATGCAAT GTTCTTAGGA TCCTCCTCCC GTCCTTCTCA TCCTTCATGT	120
	CCAGGGTGCT GCAGAAGGAC GCGGAGCAGG AGTCACAGAT GAGAGCGGAG ATCCAGGACA	180
50	TGAAGCAGGA GCTCTCCACA GTCAACATGA TGGACGAGTT TGCCAGATAT GCCAGGCTGG	240
50	AAAGAAAGAT CAACAAGATG ACGGATAAGC TCAAAACCCA TGTGAAAGCT CGGACAGCTC	300
	AATTAGCCAA GATAAAATGG GTGATAAGTG TCGCTTTCTA CGTATTGCAG GCTGCCCTGA	360
55	TGATCTCACT CATTTGGAAG TATTATTCTG TCCCTGTGGC TGTCGTGCCG AGTAAATGGA	420
	TAACCCCTCT AGACCGCCTG GTAGCCTTTC CTACTAGAGT AGCAGGTGGT GTTGGAATTA	480
	CCTGTTGGAT TTTAGTCTGT AACAAAGTTG TCGCTATTGT GCTTCATCCG TTCAGCTGAA	540

	CAGGAGGATG GATACAGCCG CGAGGCTAAA AAACGGATTT CCTCTTCCTA GCTTAAAATC	600
	TGATTTACAC TGTTTTGTTT TTTAAGAAAC AAAAGTGCAT AGTTTAGATT TTTTTTTTTG	660
5	TTGAATATGT TTGTTCTTGG ACTTTATGAG AGAGTCTTAT AAGAATCACG ATTTTCTACA	720
	CCTGTCATTG AGCCAAGAAA GTCCAGTTTA TGACACGTAT GTACTAGTGA ACACCGTCCT	780
10	CGATCTGTAC GAAATGTGAA ATGTTTAGGG ACATCTCCAT GCTGTCACTT GTGATTTGCC	840
10	CTCTTATGTA TTTTGGTCAT ATTGCCAACT GGAAAGTCAA AATTTTCTAA CAACTTTAAG	900
	TAAGTTCTTT GAAGACTTAG TGCTGTTTTT AATCCAGTTT AGAAAGTAAC TTAATTTTAA	960
15	TACCACTACT AAAAATTCGA AAATTTCTTC TTTAATCACA TTCAATATGG TTAAAAGAAC	1020
	AACACTAATT GACATTGCGT GGGCTTTTTC TCCCTTTGTT TAAAATGTCA TTTGTTGAGC	1080
20	AAGAGTTGTA TAGTATTATC TACTTACTTG AGGCTGTTAA TTTTTCATTA CAGTGTTTTG	1140
20	TAAATGTATC CACGAGACCA TGATGCATTG TTTTGTGCTC AACTTGTGTT TTGTATTTAA	1200
	AGCATTTTGA ATGAAGTGTA TTTTATAAGC ATTTAATATT TATGCTCTTT AGAATGGAAC	1260
25	ACAGAAAACA AACCTTATAA GTCCTGATTA ATCTGAACCA ATAACCTGTG TGGCCTACAA	1320
	AGTATAATTC TATTAAATGT TCCTTAAAAC AAAAAAAAA AAAAAAAAA AAAA	1374
30		
	(2) INFORMATION FOR SEQ ID NO: 47:	•
35	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 596 base pairs	
	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
	(D) TOPOLOGY: linear	
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 47:	
	GAATTCGNCA CGAGATTACT TGGACATGAA AGAACTCAGG TTCAAGTTTA TTCATTTACT	60
45	AAGTTAGTTA AATCATGTGC CTTCCATGAG CCTTCATTTG GTAACTTGGA AAATGGAAAT	120
	AATAACACTA GTCATATATA TTCTACACTG CTACCATATG GACCAAAGGG ATTATAGATT	180
	ACAATCACCA TCATTCCTGC TGACAGGTAT ATAGAAAACA ATTTCATTGA AGAAAAGTCC	240
50	TTACATTTAT CCTTTTCCTA ATATCTGCAT GGGTAAACTA ATAAATATAG TCATTAGAAA	300
		260
	ACCCTTATTA TTATTATTAG TTCAATGTGA GAACTGCTGC AGAAAAAATA TGCTTTATAA	360
55	ACCCTTATTA TTATTATTAG TTCAATGTGA GAACTGCTGC AGAAAAAATA TGCTTTATAA TATTTTCTTG AATATACATA ATATTCATAA ATTTTCAAAT CATTGAAAAT TACCTTAAAA	420
55		
55	TATTTTCTTG AATATACATA ATATTCATAA ATTTTCAAAT CATTGAAAAT TACCTTAAAA	420

3	(2) INFORMATION FOR SEQ ID NO. 46.	
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 851 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 48:	
15	CACATGAAGA CACACAGTGG TGAGAAGCCC TTCCGCTGCG CCCGCTGTCC TTATGCCTCT	60
	CCTCATCTGG ATAACCTGAA ACGGCACCAG CGCGTCCATA CAGGAGAGAA GCCCTACAAG	120
20	TGCCCCCTCT GCCCTTATGC CTGTGGCAAT CTGGCCAACC TCAAGCGTCA TGGTCGCATC	180
20	CACTCTGGTG ACAAACCTTT TCGGTGTAGC CTTTGCAACT ACAGCTGCAA CCAGAGCATG	240
	AACCTCAAAC GTCACATGCT GCGGCACACA GGCGAGAAGC CTTCCGCTGT GCCACCTGCG	300
25	CCTATACCAC GGGCCACTGG GACAACTACA AGCGCCACCA GAAGGTGCAT GGCCACGGTG	360
	GGGCAGGAGG GCCTGGTCTC TCTGCCTCTG AGGGCTGGGC CCCACCTCAT AGCCCACCCT	420
20	CTGTTTTGAG CTCTCGGGGC CCACCAGCCC TGGGGACTGC TGGCAGCCGG GCTGTCCACA	480
30	CAGACTCATC CTGAACTAGG TCCTTCTTCC CCATGTTTTA TACAGACGGA CCAGAAGCCA	540
	CCTTTTTCTC CCCCGCTGGC CAGGGGCTCC ACACAGACTA ACGTAGGCAC TATAAGGACC	600
35	AGCCCAACCC CATGGGCGGG GGGGCCCATA TGGACCAGGG GACCTTGCCT TGACTGAGGC	660
	ACTTCACGAG CTCAGTGAGA AGGGCCCTGT ATTCACCTCC ACTGCCCCCA GGGGCTGTGG	720
40	ACAAACCGGC TGGGGGACTG CCCAGCCTCC CACCTGTTTA TTTAACTTAT TTCAGTGCTT	780
40	TATAATAAAG GAAACACTAA CAAAGCCATG TCTATGCTGA ATTGGCAATG GCAGGCAATT	840
	TGGCCTTACC C	851
45		
	(2) INFORMATION FOR SEQ ID NO: 49:	
50	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 2020 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
55	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 49:	
	GTGAAATGAA AACAGTCTTT TTATAGCCTT TAGCTTGTGA GTTTGGAAGT TTGGGGGGTC	60
60	TENDETETTET THE COUNTY COUNTY CASTAGE GAGGAGAGTT GAGGCTTTTC TRACGTCCAT	120

	ACACAGACCC	AGGTGAACAC	GCTGACTGTG	AACCTGCCCT	GTATCCGGAG	CTGTGCTGGG	180
5	CACTGAGGGG	ATGCAACAAA	ATTAGGAGAG	GWTCCTTGCT	CCCAACGTCT	ACTTCTCCTA	240
	CCTCAACAGG	GGTCCAGGGT	GCAGTGAACT	CAGTTCTTGG	CCCTTGGGTG	AGGATTCATG	300
	GATGAATGAA	AGCTAGACCT	GATGGGGAGG	CATTATGACT	AAATAGGCCC	AGCCTCCTTC	360
10	CCTTCCAGCT	CTGTCCTAGG	AGCATAGGCG	GGAAATCTGA	GTAGAGTCTG	ACTGCAGTTT	420
	TTGCTTATGA	TTTGTAAAAG	CCGTCATGGG	GTCAATAAGA	AAATAGGGGT	GATGGAGGGG	480
15	GAGAAGCCCA	GGACTGGGAG	AATCGCACGT	GCCCCAGGGG	TTTTCACCAA	GGATTTTCAA	540
	GACAAACTGG	AGTAAGAATT	AAAGCCCCAG	AGGATTTAAT	TATCCTGGTT	TGCAAAAGAG	600
	CCTCCCATGC	CAGTACCGCC	CAGCCTTGGA	GGCCGGAATG	CTCATGGCCC	CTGTGGTCTG	660
20	CTTGTCCTTC	AGCCCATGCC	CAGCAGATAC	CTCTCTGACT	GGAGACGGC	TCAAAGCTGG	720
	ATTAGAAAGG	GGAGMGGCAC	TTGTGACTTT	GTTTGACTCT	GTGACTCACT	TCCTCGCTCA	780
25	CACCTTGTTT	GAACTACTGG	ACTTTCAACT	GGCTTTCCTT	AGGTCAGGCA	AGCAGACAGC	840
	TCCCCACTGA	AGAGGTCTGT	ACAGTGACAA	CCCGGGCCGG	CAGCAAGGAC	ACAGATGCAG	900
	CCACAGTAAG	GCTCCATCAG	GACTGGGTCA	GTGATGGCAA	CAGGATGGCC	AAGGATGGCT	960
30	CTAGAACAYT	CTGTCCATGC	GTCACTCCCC	CCAGTTTTRT	TTTTAGCTTT	GGCTTCAGGG	1020
	AGTGACAGCC	ATCACAAATA	GCCACATTCT	GCTCTACTCT	CCAACATACC	AGATTSTACA	1080
35	CTGTTGTTAT	TTCATGAGAC	GTGAATGTTG	CAGAGAGTGG	GGGGATTCTG	GTTGTTAAGG	1140
	AACTTACACT	GGGGAGCTTT	ACTCTTCCGT	GTCAACAATG	TGACTACATG	TTCTCCAGAT	1200
	TAGCCACACA	TGCAAACATC	AGTGTCCTTĆ	TAGCTTTANC	CGAGAAAGAA	ACCAGTCCCA	1260
40	GGGAATGAAT	GGTGGTCTCC	CCACTCCCGG	CAGCACTTTA	GGCAGCCCAT	AAGCTATGCG	1320
	AGAATGTGAA	CGCTCACCTT	GCTCCGTCAC	GGTTCTGACC	TACCACATAA	ACAGGAAGAA	1380
45	GCCAGTGACC	GGAACAGCTC	TAGGAATAAC	AAGTCAGAAT	AGAAGTGTCC	TTTATATTAC	1440
	CAGAAAATAT	GGGCTTGGCC	TAAGTCGCTG	TCTCCTAACC	TGCCGGGGTC	ATTCCCCACC	1500
	AAACACCCCA	TACTAAGGAG	CCATGAGCCA	CCTGGACATT	CACCTTTTCT	TTGACCATCT	1560
50	GGAGTCTGGG	GCAACTTAAG	GAAGGCNCCA	CACAGTGGTG	CAGGCACATT	TCCAAGCGTA	1620
	GGTGTCCCTG	GCTTTTGTGG	CCAAAGCTAG	TGTTATGGTC	AACAACAGGC	CAGGGTCTGT	1680
55	GGGGCACTGA	CCTTGAAAGT	GGCAAAATGG	AGGTTTCACA	GGCTGTGCGG	GAGCAGGACG	1740
	GCTTGCTTCA	TCTAACAATC	TCAGTTTCCT	TTAAAAAAAG	AAAGAAAGGA	AAAGATTTCA	1800
	TAAGCAGGTG	TCAGTGGACA	GTTTAAGYAC	TTAACCATTT	CTCTTTCTTC	TTATGGATGT	1860
60	GAACTGTGCT	GTGGATAAAT	CATTTGTATT	TCTTGAATGT	TCTCTATGAC	TAACAGTTAT	1920

	TAAGTCGGTT GTGTATATGT GTAACTAATG TAACTGCCTT TTAAAATTTC ATTACAATAA	1980
5	AAATGACTTT GCTCTGAAMA AAAAAAAAAA AAAAACTCGA	2020
10	(2) INFORMATION FOR SEQ ID NO: 50: (i) SEQUENCE CHARACTERISTICS:	
15	(A) LENGTH: 2432 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 50:	
20	ATGAAGGGTC GTTGGTGGGA AAGATGGCGG CGACTCTGGG ACCCCTTGGG TCGTGGCAGC	60
20	AGTGGCGGCG ATGTTTGTCG GCTCGGGATG GGTCCAGGAT GTTACTCCTT CTTCTTTGT	120
	TGGGGTCTGG GCAGGGGCCA CAGCAAGTCG GGGCGGGTCA AACGTTCGAG TACTTGAAAC	180
25	GGGAGCACTC GCTGTCGAAG CCCTACCAGG GTGTGGGCAC AGGCAGTTCC TCACTGTGGA	240
	ATCTGATGGG CAATGCCATG GTGATGACCC AGTATATCCG CCTTACCCCA GATATGCAAA	300
•	GTAAACAGGG TGCCTTGTGG AACCGGGTGC CATGTTTCCT GAGAGACTGG GAGTTGCAGG	360
30	TGCACTTCAA AATCCATGGA CAAGGAAAGA AGAATCTGCA TGGGGATGGC TTGGCAATCT	420
	GGTACACAAG GAATCGGATG CAGCCAGGGC CTGTGTTTGG AAACATGGAC AAATTTGTGG	480
35	GGCTGGGAGT ATTTGTAGAC ACCTACCCCA ATGAGGAGAA GCAGCAAGAG CGGGTATTCC	540
	CCTACATCTC AGCCATGGTG AACAACGGCT CCCTCAGCTA TGATCATGAG CGGGATGGGC	600
	GGCCTACAGA GCTGGGAGGC TGCACAGCCA TTGTCCGCAA TCTTCATTAC GACACCTTCC	660
40	TGGTGATTCG CTACGTCAAG AGGCATTTGA CGATAATGAT GGATATTGAT GGCAAGCATG	720
	AGTGGAGGGA CTGCATTGAA GTGCCCGGAG TCCGCCTGCC CCGCGGCTAC TACTTCGGCA	780
45	CCTCCTCCAT CACTGGGGAT CTCTCAGATA ATCATGATGT CATTTCCTTG AAGTTGTTTG	840
	AACTGACAGT GGAGAGAACC CCAGAAGAGG AAAAGCTCCA TCGAGATGTG TTCTTGCCCT	900
	CAGTGGACAA TATGAAGCTG CCTGAGATGA CAGCTCCACT GCCGCCCCTG AGTGGCCTGG	960
50	CCCTCTTCCT CATCGTCTTT TTCTCCCTGG TGTTTTCTGT ATTTGCCATA GTCATTGGTA	1020
	TCATACTCTA CAACAAATGG CAGGAACAGA GCCGAAAGCG CTTCTACTGA GCCCTCCTGC	1080
55	TGCCACCACT TTTGTGACTG TCACCCATGA GGTATGGAAG GAGCAGGCAC TGGCCTGAGC	1140
	ATGCAGCCTG GAGAGTGTTC TTGTCTCTAG CAGCTGGTTG GGGACTATAT TCTGTCACTG	1200
60	GAGTTTTGAA TGCAGGGACC CCGCATTCCC ATGGTTGTGC ATGGGGACAT CTAACTCTGG	1260

	TCTGGGAAGC	CACCCACCCC	AGGGCAATGC	TGCTGTGATG	TGCCTTTCCC	TGCAGTCCTT	1320
	CCATGTGGGA	GCAGAGGTGT	GAAGAGAATT	TACGTGGTTG	TGATGCCAAA	ATCACAGAAC	1380
5	AGAATTTCAT	AGCCCAGGCT	GCCGTGTTGT	TTGACTCAGA	AGGCCCTTCT	ACTTCAGTTT	1440
	TGAATCCACA	AAGAATTAAA	AACTGGTAAC	ACCACAGGCT	TTCTGACCAT	CCATTCGTTG	1500
10	GGTTTTGCAT	TTGACCCAAC	CCTCTGCCTA	CCTGAGGAGC	TTTCTTTGGA	AACCAGGATG	1560
	GAAACTTCTT	CCCTGCCTTA	CCTTCCTTTC	ACTCCATTCA	TTGTCCTCTC	TGTGTGCAAC	1620
	CTGAGCTGGG	AAAGGCATTT	GGATGCCTCT	CTGTTGGGGC	CTGGGGCTGC	AGAACACACC	1680
15	TGCGTTTCAC	TGGCCTTCAT	TAGGTGGCCC	TAGGGAGATG	GCTTTCTGCT	TTGGATCACT	1740
	GTTCCCTAGC	ATGGGTCTTG	GGTCTATTGG	CATGTCCATG	GCCTTCCCAA	TCAAGTCTCT	1800
20	TCAGGCCCTC	AGTGAAGTTT	GGCTAAAGGT	TGGTGTAAAA	ATCAAGAGAA	GCCTGGAAGA	1860
	CATCATGGAT	GCCATGGATT	AGCTGTGCAA	CTGACCAGCT	CCAGGTTTGA	TCAAACCAAA	1920
	AGCAACATTT	GTCATGTGGT	CTGACCATGT	GGAGATGTTT	CTGGACTTGC	TAGAGCCTGC	1980
25	TTAGCTGCAT	GTTTTGTAGT	TACGATTTTT	GGAATCCCAC	TTTGAGTGCT	GAAAGTGTAA	2040
	GGAAGCTTTC	TTCTTACACC	TTGGGCTTGG	ATATTGCCCA	GAGAAGAAAT	TTGGCTTTTT	2100
30	TTTTCTTAAT	GGACAAGAGA	CAGTTGCTGT	TCTCATGTTC	CAAGTCTGAG	AGCAACAGAC	2160
	CCTCATCATC	TGTGCCTGGA	AGAGTTCACT	GTCATTGAGC	AGCACAGCCT	GAGTGCTGGC	2220
	CTCTGTCAAC	CCTTATTCCA	CTGCCTTATT	TGACAAGGGG	TTACATGCTG	CTCACCTTAC	2280
35	TGCCCTGGGA	TTAAATCAGT	TACAGGCCAG	AGTCTCCTTG	GAGGGCCTGG	AACTCTGAGT	2340
	CCTCCTATGA	ACCTCTGTAG	CCTAAATGAA	ATTCTTAAAA	TCACCGATGG	AACCAAAAAA	2400
40	AAAAAAAAA	AAAAAAAA	AAAAAAAA	AA			2432
	(2) INFORMA	TION FOR SE	EQ ID NO: 51	L:			
45	(i)	SEQUENCE CI	- HARACTERIST:	ICS:			
	, -,	(A) LEN	GTH: 2340 b	ase pairs			
50		(C) STR	ANDEDNESS:	double			
50	4.4.		OLOGY: line				
			DESCRIPTION	_			
55	GACGCTGGGG	GCGGGTGGGG	GCGCGGGGTA	CCGGGCTGGA	CGGCCGGCCG	GCGCCCCTC	60
	ATTAGTATGC	GGACGAAGCG	GCGGGCTGCG	CGGAGNGACG	TCCCCTGCAG	CCGCGGACCG	120
	AGGCAGCGGC	GGCACCTGCC	GGCCGAGCAA	TGCCAAGTGA	GTACACCTAT	GTRAAACTGA	180
60	GAAGTGATTG	СПССВССССТ	TCCCTCC A AT	CCTACACCCC	3CCTC333CC	3 3 C 3 TC 3 C 3 3	240

	GGCCCAGCTT GTTATTAAAA GACATCCTCA AATGTACATT GCTTGTGTTT GGAGTGTGGA	300
5	TCCTTTATAT CCTCAAGTTA AATTATACTA CTGAAGAATG TGACATGAAA AAAATGCATT	360
3	ATGTGGACCC TGACCATGTA AAGAGAGCTC AGAAATATGC TCAGCAAGTC TTGCAGAAGG	420
	AATGTCGTCC CAAGTTTGCC AAGACATCAA TGGCGCTGTT ATTTGAGCAC AGGTATAGCG	480
10	TGGACTTACT CCCTTTTGTG CAGAAGGSCC CCAAAGACAG TGAAGCTGAG TCCAAGTACG	540
	ATCCTCCTTT TGGGTTCCGG AAGTTCTCCA GTAAAGTCCA GACCCTCTTG GAACTCTTGC	600
15	CAGAGCACGA CCTCCCTGAA CACTTGAAAG CCAAGACCTG TCGGCGCTGT GTGGTTATTG	660
13	GAAGCGGAGG AATACTGCAC GGATTAGAAC TGGGCCACAC CCTGAACCAG TTCGATGTTG	720
	TGATAAGGTT AAACAGTGCA CCAGTTGAGG GATATTCAGA ACATGTTGGA AATAAAACTA	780
20	CTATAAGGAT GACTTATCCA GAGGGCGCAC CACTGTCTGA CCTTGAATAT TATTCCAATG	840
	ACTIATTIGT TGCTGTTTTA TTTAAGAGTG TTGATTTCAA CTGGCTTCAA GCAATGGTAA	900
25	AAAAGGAAAC CCIGCCATTC TGGGTACGAC TCTTCTTTTG GAAGCAGGTG GCAGAAAAAA	960
<i>23</i>	TCCCACTGCA GCCAAAACAT TTCAGGATTT TGAATCCAGT TATCATCAAA GAGACTGCCT	1020
	TTGRACATCC TTCAGTACTC AGAGCCTCAG TCAAGGTTCT GGGGGCCGAG ATAAGAACGT	1080
30	CCCCACAATC GGTGTCATTG CCGTTGTCTT AGCCACACAT CTGTGCGATG AAGTCAGTTT	1140
	GGCGGGTTTT GGATATGACC TCAATCAACC CAGAACACCT TTGCACTACT TCGACAGTCA	1200
35	ATGCATGGCT GCTATGAACT TTCAGACCAT GCATAATGTG ACAACGGAAA CCAAGTTCCT	1260
33	CTTAAAGCTG GTCAAAGAGG GAGTGGTGAA AGATCTCAGT GGAGGCATTG ATCGTGAATT	1320
	TTGAACACAG AAAACCTCAG TTGAAAATGC AACTCTAACT CTGAGAGCTG TTTTTGACAG	1380
40	CCTTCTTGAT GTATFTCTCC ATCCTGCAGA TACTTTGAAG TGCAGCTCAT GTTTTTAACT	1440
	TTTAATTTAA AAACACAAAA AAAATTTTAG CTCTTCCCAC TTTTTTTTTC CTATTTATTT	1500
45	GAGGTCAGTG TITGTTTTTG CACACCATTT TGTAAATGAA ACTTAAGAAT TGAATTGGAA	1560
73	AGACTTCTCA AAGAGAATTG TATGTAACGA TGTTGTWITG ATTTTTAAGA AAGTAATTTA	1620
	ATTTGTAAAA CTTCTGCTCG TTTACACTGC ACATTGAATA CAGGTAACTA ATTGGAAGGA	1680
50	GAGGGGAGGT CACTCTTTTG ATGGTGGCCC TGAACCTCAT TCTGGTTCCC TGCTGCGCTG	1740
	CTTGGTGTGA CCCACGGAGG ATCCACTCCC AGGATGACGT GCTCCGTAGC TCTGCTGCTG	1800
55	ATACTGGGTC TGCGATGCAG CGGCGTGAGG CCTGGGCTGG TTGGAGAAGG TCACAACCCT	1860
JJ	TCTCTGTTGG TCTGCCTTCT GCTGAAAGAC TCGAGAACCA ACCAGGGAAG CTGTCCTGGA	1920
	GGTCCCTGGT CGGAGAGGGA CATAGAATCT GTGACCTCTG ACAACTGTGA AGCCACCCTG	1980
60	GGCTACAGAA ACCACAGTCT TCCCAGCAAT TATTACAATT CTTGAATTCC TTGGGGATTT	2040

	TTTACTGCCC TTTCAAAGCA CTTAAGTGTT AGATCTAACG TGTTCCAGTG TCTGTCTGAG	2100
5	GTGACTTAAA AAATCAGAAC AAAACTTCTA TTATCCAGAG TCATGGGAGA GTACACCCTT	2160
J	TCCAGGAATA ATGTTTTGGG AAACACTGAA ATGAAATCTT CCCAGTATTA TAAATTGTGT	2220
	ATTTAAAAAA AAGAAACTTT TCTGAATGCC TACTGGCGGT GTATACCAGG CAGTGTGCCA	2280
10	GTTTAAAAAG ATGAAAAAGA ATAAAAACTT TTGAGGAAMA AAAAAAAAAA AAAAACTCGA	2340
15	(2) INFORMATION FOR SEQ ID NO: 52:	
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 601 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 52:	
25	AGTAGGGGAG ACTGAGACTG ACCGGTAGCC AGGCAGGCGG ACGACGCACG CCCGGACAGA	60
	CTGAGCAGGC GCCGGAGAAC CACTCACAGG TTCCCCCCGC CTTTCCCTTT GAAANCTAGG	120
30	CTTTTGCCTT TCCCGTGGCG CCCGAGAGAG AATGCTGGAC TCTGCCGACT TCAGCGCAAC	180
	TAANGATTTC TCAAGCTAGG GGACAAACGA TCAGCCCAAT CCTGAGAAGG GGGGAACCAA	240
	GCACCCCGTC CCCATCCCCC TCCCCTCCCC CGACTAAACT CGGGCGCCAA ACCCAGCCCT	300
35	TCTCTAACCA CCCTACTTCC TCCTCCTT TCTAGCATGG TGGCTGTATG GACAGTCTGA	360
	CAGAACAGAG ACTGACATCT CCCAATCTGC CGGCCCCCCA CCTGGAACAC TACAGTGTTC	420
40	TGCATTGCAC CATGACCCTG GATGTGCAAA CTGTAGTCGT TTTTGCCGTG ATTGTAGTCC	480
	TCCTGCTTGT CAATGTCATA CTCATGTTTT TCCTGGGAAC GCGCTGAATG GAGTCCAGNC	540
	ACCTGAGCTG TCGCGAACTC TCGCTTTGAT TTCATCCCGA GAGCCACCGA GAAGAAAAAA	600
45	A	601
50	(2) INFORMATION FOR SEQ ID NO: 53:	
	(i) SEQUENCE CHARACTERISTICS:	
55	(A) LENGTH: 359 base pairs (B) TYPE: nucleic acid	
55	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 53:	
60	CTCGTGCCGA ATTCGGCACG AGAGATGGTA CTTTTAAGAG GTAATTAGGT TGCTAAGATG	60

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	GATTAACATC TTTCTCTTGA CACTGAGACT GGGTTCTCCT GGGAATGGTT AGTTCCCAAG	120
_	AGAGTGAGTT GTTATAAAAC AATGCTGCCT CTTCTATTTT GCGCTTTTTG TTTGCACAAA	180
5	CTCGGTCCCC TTCTGTTTCT CTACGATGTT TTGATGCRGC ATGAGGCAGT CATGAGAACC	240
	CACCAGATAC AGCTGCCTGA TCCTGAATTT CCCAGCCAAC AGAACCAAGT GCTAAATAAA	300
10	ACTCTTTTTA ATAAGTTAAA AAAAAAAAAA AAAAAAAAA AANAAANANA AAAAAA	359
15	(2) INFORMATION FOR SEQ ID NO: 54:	
	(i) SEQUENCE CHARACTERISTICS:	
20	(A) LENGTH: 1141 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 54:	
25	GGCACGAGCT GCTGAGGCGT GAGAATGGCG TCCCGCGGCC GGCGTCCGGA GCATGGCGGA	60
	CCCCCAGAGC TGTTTTATGA CGAGACAGAA GCCCGGAAAT ACGTTCGCAA CTCACGGATG	120
	ATTGATATCC AGACCAGGAT GGCTGGGCGA GCATTGGAGC TTCTTTATCT GCCAGAGAAT	180
30	AAGCCCTGTT ACCTGCTGGA TATTGGCTGT GGCACTGGGC TGAGTGGAAG TTATCTGTCA	240
	GATGAAGGGC ACTATTGGGT GGGCCTGGAT ATCAGCCCTG CCATGCTGGA TGAGGCTGTG	300
35	GACCGAGAGA TAGAGGGAGA CCTGCTGCTG GGGGATATGG GCCAGGGCAT CCCATTCAAG	360
	CCAGGCACAT TTGATGGTTG CATCAGCATT TCTGCTGTGC AGTGGCTCTG TAATGCTAAC	420
	AAGAAGTCTG AAAACCCTGC CAAGCGCCTG TACTGCTTTT TTGCTTCTCT TTTTTCTGTT	480
40	CTCGTCCGGG GATCCCGAGC TGTCCTGCAG CTGTACCCTG AGAACTCAGA GCAGTTGGAG	540
	CTGATCACAA CCCAGGCCAC AAAGGCAGGC TTCTCCGGTG GCATGGTGGT AGACTACCCT	600
45	AACAGTGCCA AAGCAAAGAA ATTCTACCTC TGCTTGTTTT CTGGGCCTTC GACCTTTATA	660
	CCAGAGGGC TGAGTGAAAA TCAGGATGAA GTTGAACCCA GGGAGTCTGT GTTCACCAAT	720
	GAGAGGTTCC CATTAAGGAT GTCGAGGCGG GGAATGGTGA GGAAGAGTCG GGCATGGGTG	780
50	CTGGAGAAGA AGGAGCGGCA CAGGCGCCAG GGCAGGGAAG TCAGACCTGA CACCCAGTAC	840
	ACCGGCCGCA AGCGCAAGCC CCGCTTCTAA GTCACCACGC GGTTCTGGAA AGGCACTTGC	900
55	CTCTGCACTT TTCTATATTG TTCAGCTGAC AAAGTAGTAT TTTAGAAAAG TTCTAAAGTT	960
	ATAAAAATGT TTTCTGCAGT AAAAAAAAAG TTCTCTGGGC CGGGCGTGGT GGCTCACACC	1020

TGTAATCCCA GCACCTTGGG AGGCTGAGGT GGGAGGATCA TTTGAGGCCA GGAGTTTGAG

	ACCTGCCTGG GCAACATAAT GAAACTTCCT TTCCAGGGAG AAAAAAAAAA	1140
	A	1141
5		
	(2) INFORMATION FOR SEQ ID NO: 55:	
10		
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 1560 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 55:	
	TCCTTCTCTG GGGCGGTCGC GTTGGCAGCG GATGCGGGAA GCCGGACTCT GGGCGTCATG	-
20		60
20	TACTACAAGT TTAGTGGCTT CACGCAGAAG TTGGCAGGAG CATGGGCTTC GGAGGCCTAT	120
	AGCCCGCAGA TINAAAGCCT GTGGTTTCCA CAGAAGCACC ACCTATCATA TITGCCACAC	180
25	CAACTAAACT GACCTCCGAT TCCACAGTGT ATGATTATGC TGGGAAAAAC AAAGTTCCAG	240
	AGCTACAAAA GTTTTTCCAG AAAGCTGATG GTGTGCCCGT CTACCTGAAA CGAGGCCTGC	300
	CTGACCAAAT GCTTTACCGG ACCACCATGG CGCTGACTGT GGGAGGGACC ATCTACTGCC	360
30	TGATCGCCCT CTACATGGCT TCGCAGCCCA AAAACAAATG AGTTAGGCTG CAGAGGACTG	420
	GTTTGTTTTT TGGCATAAAC CCTTTGAAGT TCCTTTTTCA TTGTTAAATT AAAATTTTTT	480
35	TTTTTACTTG GATGGCTTAA CATTTTTGCA AGAAAAATAG GAAGATATGA AGATGATGTT	540
55	TTGGTTTGTT TATGAAATGC ATATGGCTTG TCAGAGCTCA TTCGACAGTT AAAGCCATTG	600
	TTTAAAGAAA CGGTGCTTTG CTCTGTGTTT GTGCTCCTGA TTTCCCTGGA GGTTCTGGAT	660
40	GAAGGCTGAA CACAGGCTTG TTAATGTCAG TCTGTGCTGA GGACCTCAGG GACTTGAGGT	720
	TGCATTTTTG AGCATGGGGT GCAGGAGCCT TTCTGGATTT GGATGTGGCT ATGGAAAGAA	780
	CACAGAAGCC AAGGTCATGT GCATGAAATG AGGAGTTTGA GTTAGTCACC TCGGGGATTT	840
45	TTTCCATTTT GCAGTAAAAT GTTAAATTAA TGTAGCCTGC CTCTATTTGT TGGGCAGGTA	900
	ATTTCAAAGG GTTATTTGCC TCATCTCCTA TCTTTAGTGA AATCTTATGT GTAATTGTGT	960
50	GTATTTATTC CACCGTGGGA ACAGAGAATA CCTGTTTAGT GTTGCACTTT AGACTGGTGT	
		1020
	CTGTTTTGTT AATGCAGCTG TGCCACAAAT TCTCCTTTAT CTTTTAAAAA TGTTATAGCT	1080
55	TTAAATTTTG ATTTATTTTG ACTGTGGAAT AAATACATGA ATGAAAAATT TTAAGTTTGA	1140
	AGTTCTTTGA ATGACCTTTC AGAGTAATTT CAGAACACCA GCAGCATCTT AAACCTGAGT	1200
	CTAATTTCTT TCTTGTTAAT TAGGCACCAG ATAATCTTTA TAAAAATGGTC TTAAAAGCTA	1260
60	GTAATAGGAG CTTAATGGCA ATKGATGATT ACCACAKGGT TTTTTATAAA AACCTGCCTG	1320

	CCCCTWAGTG AAAGGTACCT GTAACYCACA GTYCATTTAG ACACTAATTT CCTYTGCYGT	1380
5	CATGATTGGK AGACTTCACT TACCCTATAT TAATTTTGAA AAAAGGTGGA ATTTTATTAT	1440
3	ATATGAAGGA ATAGTTTGTA TCTTACCATA GCACAGAACA GTGACCTCTT GCTCAGGATA	1500
	AGATGTGGTG ATTTGAAAAT ACTCATAGTA GCCTTGCAGT GATACCTCTC TCNCTCTCTC	1560
10		
	(2) INFORMATION FOR SEQ ID NO: 56:	
15	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 1507 base pairs (B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double	
20	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 56:	
	GGAACGCAGA GCGGAGCGTG GAGAGCGGAG CGAAGCTGGA TAACAGGGGA CCGATGATGT	60
25	GGCGACCATC AGTTCTGCTG CTTCTGTTGC TACTGAGGCA CGGGGCCCAG GGGAAGCCAT	120
	CCCCAGACGC AGGCCCTCAT GGCCAGGGGA GGGTGCACCA GGCGGCCCCC CTGAGCGACG	180
30	CTCCCCATGA TGACGCCCAC GGGAACTTCC AGTACGACCA TGAGGCTTTC CTGGGACGGG	240
30	AAGTGGCCAA GGAATTCGAC CAACTCACCC CAGAGGAAAG CCAGGCCCGT CTGGGGCGGA	300
	TCGTGGACCG CATGGACCGC GCGGGGGACG GCGACGGCTG GGTGTCGCTG GCCGAGCTTC	360
35	GCGCGTGGAT CGCGCACACG CAGCAGCGGC ACATACGGGA CTCGGTGAGC GCGGCCTGGG	420
	ACACGTACGA CACGGACCGC GACGGGCGTG TGGGTTGGGA GGAGCTGCGC AACGCCACCT	480
40	ATGGCCACTA CGCGCCCGGT GAAGAATTTC ATGACGTGGA GGATGCAGAG ACCTACAAAA	540
40	AGATGCTGGC TCGGGACGAG CGGCGTTTCC GGGTGGCCGA CCAGGATGGG GACTCGATGG	600
	CCACTCGAGA GGAGCTGACA GCCTTCCTGC ACCCCGAGGA GTTCCCTCAC ATGCGGGACA	660
45	TCGTGATTGC TGAAACCCTG GAGGACCTGG ACAGAAACAA AGATGGCTAT GTCCAGGTGG	720
	AGGAGTACAT CGCGGATCTG TACTCAGCCG AGCCTGGGGA GGAGGAGCCG GCGTGGGTGC	780
50	AGACGGAGAG GCAGCAGTTC CGGGACTTCC GGGATCTGAA CAAGGATGGG CACCTGGATG	840
	GGAGTGAGGT GGGCCACTGG GTGCTGCCCC CTGCCCAGGA CCAGCCCCTG GTGGAAGCCA	900
	ACCACCTGCT GCACGARAGC GACACGGACA AGGAYGGGCG GCTGAGCAAA GCGSAAATCC	960
55	TGGGTAATTG GAACATGTTT GTGGGCAGTC AGGCCACCAA CTATGGYGAG GACCTGACCC	1020
	GGCACCACGA TGAGCTGTGA GCMCCGNGCA CCTGCCACAG CCTCAGAGGC CCGCACAATG	1080
60	ACCGGAGGAG GGGCCGCTGT GGTCTGGCCC CCTCCCTGTC CAGGCCCCGC AGGAGGCAGA	1140

	TGCAGICCCA GGCAICCICC TRCCCCIGGG CICICAGGGA CCCCCCIGGGT CGGCTTCTGT	1200
	CCCTGTCACA CCCCCAACCC CAGGGAGGGG CTGTCATAGT CCCAGAGGAT AAGCAATACC	1260
5	TATTTCTGAC TGAGTCTCCC AGCCCAGACC CAGGGACCCT NGGCCCCAAG CTCAGCTCTA	1320
	AGAACCGCCC CAACCCCTCC AGCTCCAAAT CTGAGCCTCC ACCACATAGA CTGAAACTCC	1380
10	CCTGGCCCCA GCCCTCTCCT GCCTGGCCTG GCCTGGGACA CCTCCTCTCT GCCAGGAGGC	1440
10	AATAAAAGCC AGCGCCGGGA AAAAAAAAAA AAAAAAAAA AAAAAAAA	1500
	AAAAAAN	1507
15		
	(2) INFORMATION FOR SEQ ID NO: 57:	
20	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 450 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 57:	
	GAATTCGGCA CGAGCAGTGT CCAACACTGT AGCTGGTGCC TGCCAGGTTC CCAGTGGCTG	60
30	GGGTCACCAG GTCTGAAGAG AGATGTGCTG GCTGCGGGCA TGGGSCCAGA TCYTCCTGCC	120
	AGTTTTCYTC TCCYTCTTTC TCATCCAATT GCTTATCAGC TTCTCAGAGA ATGGTTTTAT	180
35	CCACAGCCCC AGGAACAATC AGAAACCAAG AGATGGGAAT RAAGAGGAAT GTGCTGTAAA	240
	GAAGAGTTGT CAATTGTGCA CAGAAGATAA GAAATATATG ATGAATAGAT AATTGAAAAG	300
	AGATCCTCCA GAAAGAGCAG AAGGAAGTTT CTTCAATGGC TTCCTTCAGG ATTTTAATCA	360
40	TCCTTACAGC CTCTTTGAGA ATGATTGAAC TTCCAAATTC CCTGAAGTTA AAATTTTAAA	420
	TTCTATTAAA CATTTTTTCG AGTAAAAAA	450
45		
	(2) INFORMATION FOR SEQ ID NO: 58:	
50	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1147 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
55	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 58:	
	GGCACGAGAC CCATTGAGCA GAAGGAGGCC AGGTGGGAAA GCTCCTGGGA AGAGCAGCCA	60
60	GACTGGACAC TGGGCTGCTT GAGTCCTGAG TCACAATTCA GAATTCCTGG GCTCCCTGGG	120

	TGCATTCTAT	CATTCCAGTT	GAAAGTTTGC	TTCCTTCCAG	TCATGTGGCT	CTTCATTCTA	180
	CTCTCCTTGG	CTCTCATTTC	AGATGCCATG	GTCATGGATG	AAAAGGTCAA	GAGAAGCTTT	240
5	GTGCTGGACA	CGGCTTCTGC	CATCTGCAAC	TACAATGCCC	ACTACAAGAA	TCACCCCAAA	300
	TACTGGTGCC	GAGGCTATTT	CCGTGACTAC	TGCAACATCA	TCGCCTTCTC	CCCTAACAGC	360
10	ACCAATCATG	TGGCCCTGAA	GGACACAGGG	AACCAGCTCA	TTGTCACTAT	GTCCTGCCTG	420
10	AACAAAGAAG	ACACGGGCTG	GTACTGGTGT	GGCATCCAGC	GGGACTTTGC	CAGGGATGAC	480
	ATGGATTTTA	CAGAGCTGAT	TGTAACTGAC	GACAAAGGAA	CCTGGCCAAT	GACTTTGGTC	540
15	TGGGAAAGAC	TATCAGGCAC	AAAACCAGAA	GCTGCAAGGC	TCCCAAAGTT	GTCCGCAAGG	600
	CTGACCGCTC	CAGGACGTCC	ATTCTCATCA	TTTGCATACT	GATCACGGGT	TTGGGAATCA	660
20	TCTCTGTAAT	CAGTCATTTG	ACCAAAAGGA	GGAGAAGTCA	AAGGAATAGA	AGGGTAGGCA	720
20	ACACTTTGAA	GCCCTTCTCG	CGTGTCCTGA	CTCCAAAGGA	AATGGCTCCT	ACTGAACAGA	780
	TGTGACTGAA	GATTTTTTA	ATTTAGTTCA	TAAAGTGATG	CTACAACAGA	ATAATCACCA	840
25	TGACAACTGG	CCCCACACCT	CAGAGACTGA	TTCTGATCTC	CCAGGAATTC	TGAAGGTCCC	900
	TCTATCCTTG	ACAACAATCA	. TTTGCAGCCA	GGTAGCAACG	GCAGTAGTCA	GAGGAGCTAT	960
20	GATAGACCAC	ACCCAAGCAA	GGCTGCCCTC	: AAATAACATO	TCAAGATCTT	AGTTCTTATG	1020
30	CATTCCATCA	GTCAGAAGTC	AAGAAGAGGT	GGAGAATCTG	GATTGGGGAC	CAGGAAATCA	1080
	CTTGTATTTT	GTTAGCCAAT	AAATTCCTAC	CCAGTGTTGA	ATGAAAAAA	AAAAAAAA A	1140
35	AAAAAA						1147

40 (2) INFORMATION FOR SEQ ID NO: 59:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 777 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double
(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 59: GGCAGAGGCT CCTCAGAAGG GCGTGGGCTC TCCAGTCTTC CACAGTCCCC ACCATGCCCT 60 50 GTTGCCTTAC CGCTGACGTA GCTCACCCAT CTTTTACTTG CCTGGCTAAG ATGCATGGCA 120 TYWCATTTCC TCCTTGTTGC ACTGCAGTCA GTCCCTCACT GCCCCCATCT CCTGGAAGAG 180 55 GAGCATAAGC TTTGCAAGGT CAGCCACTTC TCTGGGGTCA CACTAGTTAC ATCAAGACAG 240 GACTCCAGCT CATATGTGCC AGTGCAGACA CTCTTCATCC ACCTGGGGCC CTGGGCTTGG 300 GACCTGGYTC CTTGCACAGC AGARGACCCG GAGGCTGAGA GGAGCTTGCG GTTGTGTCAT 360 60

	AGICACCIGG	CCAGARGGAA	CGTGAGCCCC	TCCCAAGCTG	CAGARGGARG	GARCARGCGT	420
5	GGCTGTCAGC	ACCGAGGTAG	CAGAGAATTA	ACATTCTTGT	CAGCAGAGAA	TGAAGCAGGA	480
3	ATATAATTAA	AACTTTGCCC	TTGGAATAGC	TGATTCATTT	GAATTTTATT	CCACACGTTT	540
	GAAAGAGGAA	AGAAAATGTG	AAGACTTGCA	GCCTGGTTCT	CGCCTGGCCT	GGGCTGGCCC	600
10	AGCTGTCAGG	CCCGGTTCCT	TTCTGAGCAT	TCAGTCCACT	GATGTTGACT	GAGGGCCAGG	660
	AGAGACCCTC	AGCAGGGTAT	TACCATATCA	GCCTCCTATC	GCTGCTGGGA	GAAATTACCA	720
15	TGAATTCAGT	GGCTTAAAAC	AACACACGAG	CCTCTCTGAG	CCTACCCTGG	CTCAGGA	777
20		ATION FOR SE	_				
25		(A) LEN (B) TYP (C) STR	GTH: 1191 b E: nucleic ANDEDNESS: OLOGY: line	ase pairs acid double			
	(xi) SEQUENCE I	DESCRIPTION	: SEQ ID NO	: 60:		
30	AAGANTGATT	TTCCTTACTC	TCCAAAGCGT	CAGCATTTTG	AAGTTTCTTT	TATGAAAGTG	60
	GGGGCAAGAA	TCAGGGTGAA	AATGAGTGTA	AACAAAGCCC	ATCCTGTGGT	CAGCACCCAC	120
	TGGAGGTGGC	CAGCAGAGTG	GCCTCAGATG	TTCCTGCACC	TGGCCCAGGA	GCCCAGGACA	180
35	GAGGTCAAAT	CTAGGCCCCT	TGGTCTGGCT	GGATTCATCA	GGCAAGATTC	GAAAACAAGA	240
	AAACCTCTAG	AACAAGAAAC	AATCATGTCT	GCAGCAGATA	CGGCACTGTG	GCCCTATGGC	300
40	CATGGCAATC	GTGAGCACCA	AGAGAATGAG	TTACAGAAAT	ATCTCCAATA	CAAAGACATG	360
	CATCTCCTGG	ACAGTGGACA	GTCGCTGGGA	CACACACACA	CACTTCAAGG	CTCACACAAC	420
	CTAACAGCCT	TAAATATCTG	AAGAAACAGA	ATCACGACAT	TAAGTCAGCA	GAGGGAGAGG	480
45	TAGGCTGAAG	CAGCAGGAGG	CCAATTTTAT	ATCCCACAGA	ТТТТТТТААА	AATGACTCCC	540
	CAGCAAGGGG	TGGGGAGAAA	GCCACTGATT	TAGGAGAGTT	CTTGGCTCAG	CCAACCACTG	600
50	CGGTTATCTA	CACGTTTTAC	AAAGGCACRG	AAGTAGAGAG	GGGCTGCACT	CACGACCCTC	660
	CCCAGGGCCC	GCACAGCCAG	ACACGGTGGG	TTCTTCCTTT	TTCCCTTCTG	GCCTTGGTGG	720
	AATTCCTACC	ACGGTGGCCT	CTGCCTTTGG	GACAATGCCT	TCATGCTCAT	CCCCGGGTCA	780
55	AGGATGGAGT	CTGTTACCAT	TTTCCAGGGG	AAATTCCAAG	GACCAGCCCC	GCCTCATTAC	840
	GTTCACCCCA	CAGGAAGGTG	ATCTGGAAAG	CCTGTAAACA	CGTACTCTGG	GTGGCTGAGT	900
60	GGTGTCACCA	AGCTGCTTTT	GTGCAGGGCT	GAAGCACAGA	CAAGAGGGCA	GGCAGCTGCC	960

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	GGAGGCCTGA AGTG	GGGAGA GATCCCCGCA	GGCCTGCAGG	AGCCAGGGAG	AACCTCCAAC	1020
	TGGATCTAAA CTGT	GGGACA GCCCAGGCGT	GCCCCTCTTC	ACATGGCTCC	CAGGCTCCCT	1080
5	CAAAGCCCTT CCCA	GGCCCT GCAGGAAGAG	AGGGAGGGTG	AGGAGAGGCA	GGGAGGCAG	1140
	AGGTCGCCTG AAAG	CCTGGG CTCCGAACTC	CCTCAGCAGA	GCTTTAAAGT	G	1191
10						
	(2) INFORMATION	FOR SEQ ID NO: 6	1:			
15		JENCE CHARACTERIST (A) LENGTH: 1580 I				

(B) TYPE: nucleic acid (C) STRANDEDNESS: double

(D) TOPOLOGY: linear

20 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 61:

	CCCCGCCCCC	CGCCCACGAA	GGAAGTGGCT	GCTGCTCCGG	CGCGGACCCA	GAGCCGGTTC	60
25	GGCGCGTCGA	CTGCCCAGAG	TCCGCGGCCG	GGCGCGGGAG	GAGCCAAGCC	GCCATGGCCT	120
23	ACCACAGCTT	CCTGGTGGAG	CCCATCAGCT	GCCACGCCTG	GAACAAGGAC	CGCACCCAGA	180
	T'IGCCATCTG	CCCCAACAAC	CATGAGGTGC	ATATCTATGA	AAAGAGCGGT	GCCAAATGGA	240
30	CCAAGGTGCA	CGAGCTCAAG	GAGCACAACG	GGCAGGTGAC	AGGCATCGAC	TGGGCCCCCG	300
	AGAGTAACCG	TATTGTGACC	TGCGGCACAG	ACCGCAACGC	CTACGTGTGG	ACGCTGAAGG	360
35	GCCGCACATG	GAAGCCCACG	CTGGTCATCC	TGCGGATCAA	CCGGGCTGCC	CGCTGCGTGC	420
33	GCTGGGCCCC	CAACGAGAAC	AAGTTTGCTG	TGGGCAGCGG	CTCTCGTGTG	ATCTCCATCT	480
	GTTATTTCGA	GCAGGAGAAT	GACTGGTGGG	TTTGCAAGCA	CATCAAGAAG	CCCATCCGCT	540
40	CCACCGTCCT	CAGCCTGGAC	TGGCACCCCA	ACAATGTGCT	GCTGGCTGCC	GGCTCCTGTG	600
	ACTTCAAGTG	TCGGATCTTT	TCAGCCTACA	TCAAGGAGGT	GGAGGAACGG	CCGGCACCCA	660
45	CCCCGTGGGG	CTCCAAGATG	CCCTTTGGGG	AACTGATGTT	CGAATCCAGC	AGTAGCTGCG	720
43	GCTGGGTACA	TGGCGTCTGT	TTCTCAGCCA	GCGGGAGCCG	CGTGGCCTGG	GTAAGCCACG	780
	ACAGCACCGT	CTGCCTGGCT	GATGCCGACA	AGAAGATGGC	CGTCGCGACT	CTGGCCTCTG	840
50	AAACACTACC	ACTGCTGGCG	CTGACCTTCA	. TCACAGACAA	CAGCCTGGTG	GCAGCGGGCC	900
	ACGACTGCTT	CCCGGTGCTG	TTCACCTATG	ACGCCGCCGC	GGGGATGCTG	AGCTTCGGCG	960
55	GGCGGCTGGA	CGTTCCTAAG	CAGAGCTCGC	: AGCGTGGCTT	GACGGCCCGC	GAGCGCTTCC	1020
33	AGAACCTGG	CAAGAAGGCG	AGCTCCGAGG	GTGGCACGGC	TGCGGGCGCG	GCCTAGACT	1080
	CGCTGCACA	A GAACAGCGTC	AGCCAGATCT	CGGTGCTCAC	GCGCGCAAG	GCCAAGTGCT	1140
60	CGCAGTTCTC	G CACCACTGGC	ATGGATGGCC	GCATGAGTAT	CTGGGATGTC	AAGAGCTTGG	1200

	AGTCAGCCTT	GAAGGACCTC	AAGATCAAAT	GACCTGTGAG	GAATATGTTG	CCTTCATCCT	1260
5	AGCTGCTGGG	GAAGCGGGGA	GAGGGGTCAG	GGAGGCTAAT	GGTTGCTTTG	CTGAATGTTT	1320
5	CTGGGGTACC	AATACGAGTT	CCCATAGGGG	CTGCTCCCTC	AAAAAGGGAG	GGGACAGATG	1380
	GGGAGCTTTT	CTTACCTATT	CAAGGAATAC	GTGCCTTTTT	CTTAAATGCT	TTCATTTATT	1440
10	GAAAAAAAA	AAAAATGCCC	CCAAAGCACT	ATGCTGGTCA	TGAACTGCTT	CAAAATGTGG	1500
	AGGTAATAAA	ATGCAACTGT	GTAAAAAAA	АААААААА	AAATGACCCT	CGCGATCTAG	1560
15	AACTAGNCGG	ACGCNTGGGT					1580
20	(2) INFORMA	ATION FOR SE	EQ ID NO: 62	2:			
25	(i)	(B) TYP (C) STR	GTH: 1117 b E: nucleic ANDEDNESS:	ase pairs acid double			
23	(mi		OLOGY: line		63		
		_		: SEQ ID NO GGCTGCGCAA		cooccurace	60
30				GCGCCCGGGC			60 120
				CCTCCAGGAC			180
35				GGCTGCTGCC			240
				AGCGGCCCTG			300
				TTTCAAAGCA			360
40	TGGAGCTGTG	ATTATGGCCG	TGCGGAGGCC	AGGCTGTTTC	CTCTGTCGAG	AGGAAGCTGC	420
	GGATCTGTCC	TCCCTGAAAA	GCATGTTGGA	CCAGCTGGGC	GTCCCCCTCT	ATGCAGTGGT	480
45	AAAGGAGCAC	ATCAGGACTG	AAGTGAAGGA	TTTCCAGCCT	TATTTCAAAG	GAGAAATCTT	540
	CCTGGATGAA	AAGAAAAAGT	TCTATGGTCC	ACAAAGGCGG	AAGATGATGT	TTATGGGATT	600
50	TATCCGTCTG	GGAGTGTGGT	ACAACTTCTT	CCGAGCCTGG	AACGGAGGCT	TCTCTGGAAA	660
50	CCTGGAAGGA	GAAGGCTTCA	TCCTTGGGGG	AGTTTTCGTG	GTGGGATCAG	GAAAGCAGGG	720
	CATTCTTCTT	GAGCACCGAG	AAAAAGAATT	TGGAGACAAA	GTAAACCTAC	TTTCTGTTCT	780
55	GGAAGCTGCT	AAGATGATCA	AACCACAGAC	TTTGGCCTCA	GAGAAAAAAT	GATTGTGTGA	840
	AACTGCCCAG	CTCAGGGATA	ACCAGGGACA	TTCACCTGTG	TTCATGGGAT	GTATTGTTTC	900
60	CACTCGTGTC	CCTAAGGAGT	GAGAAACCCA	TTTATACTCT	ACTCTCAGTA	TGGATTATTA	960

	ATGTATTTTA ATATTCTGTT TAGGCCCACT AAGGCAAAAT AGCCCCAAAA CAAGACTGAC	1020
	AAAAATCTGA AAAACTAATG AGGATTATTA AGCTAAAACC TGGGAAATAG GAGGCTTWAA	1080
5	ATGACTGCCM GCTGGTGCRT GCTCACACTT GGCCCAC	1117
10	(2) INFORMATION FOR SEQ ID NO: 63:	
15	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 361 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 63:	
20	CCCACGCGTG CKGGCGCCTG GCAGCCACCG CCTGGGAGGT TACTGTAAGG CCCGCAGCTC	60
	CCGCCAGCTC CCGCGGACTS CTGCCGCCTC CTTACCATGA AGCCAGTAAG TCGTCGCACG	120
25	CTGGACTGGA TTTATTCAGT GTTGCTGCTT GCCATCGTTT TAATCTCCTG GGGCTGCATC	180
23	ATCTATGCTT CGATGGTGTC TGCAAGACGA CAGCTAAGGA AGAAATACCC AGACAAAATC	240
	TTTGGGACGA ATGAAAATTT GTAACTCTTC TGGATTTAAT TATCTGAAAA TACAGTTCTT	300
30	TCCCTCATGC TTATGTAGAT ATAAAAATAA AATTCATAAT GCAAAAAAAA AAAAAAAAA	360
	G	361
35		
	(2) INFORMATION FOR SEQ ID NO: 64:	
40	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1668 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
45	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 64:	
	GGCACGAGGT CTGCCAAGCT ATAGACCATG GCTGTGAACA CATTTGTGTG AACAGTGACG	60
50	ACTCATACAC GTGCGAGTGC TTGGAGGGAT TCCGGCTCGC TGAGGATGGG AAACGCTGCC	120
50	GAAGAAGGAT GTCTGCAAAT CAACCCACCA TGGCTGCGAA CACATTTGTG TTAATAATGG	180
	GAATTCCTAC ATCTGCAAAT GCTCAKAGGG ATTTGTTCTA GCTGAGGACG GAAGACGGTG	240
55	CAAGAAATGC ACTGAAGGCC CAATTGACCT GGTCTTTGTG ATCGATGGAT CCAAGAGTCT	300
	TGGAGAAGAG AATTTTGAGG TCGTGAAGCA GTTTGTCACT GGAATTATAG ATTCCTTGAC	360
60	AATTTCCCCC AAAGCCGCTC GAGTGGGGCT GCTCCAGTAT TCCACACAGG TCCACACAGA	420

180

	GTTCACTCTG	AGAAACTTCA	ACTCAGCCAA	AGACATGAAA	AAAGCCGTGG	CCCACATGAA	480
	ATACATGGGA	AAGGGCTCTA	TGACTGGGCT	GGCCCTGAAA	CACATGTTTG	AGAGAAGTTT	540
5	TACCCAAGGA	GAAGGGCCA	GGCCCTTTCC	ACAAGGGTGC	CCAGAGCAGC	CATTGTGTTC	600
	ACCGACGGAC	GGGCTCAGGA	TGACGTCTCC	GAGTGGGCCA	GTAAAGCCAA	GGCCAATGGT	660
10	ATCACTATGT	ATGCTGTTGG	GGTAGGAAAA	GCCATTGAGG	AGGAACTACA	AGAGATTGCC	720
•	TCTGAGCCCA	CAAACAAGCA	TCTCTTCTAT	GCCGAAGACT	TCAGCACAAT	GGATGAGATA	780
	AGTGAAAAAC	TCAAGAAAGG	CATCTGTGAA	GCTCTAGAAG	ACTCCGATGG	AAGACAGGAC	840
15	TCTCCAGCAG	GGGAACTGCC	AAAAACGGTC	CAACAGCCAA	CAGTGCAACA	CAGATATCTG	900
	TTTGAAGAAG	ACAATCTTTT	ACGGTCTACA	CAAAAGCTTT	CCCATTCAAC	AAAACCTTCA	960
20	GGAAGCCCTT	TGGAAGAAAA	ACACGATCAA	TGCAAATGTG	AAAACCTTAT	AATGTTCCAG	1020
	AACCTTGCAA	ACGAAGAAGT	AAGAAAATTA	ACACAGCGCT	TAGAAGAAAT	GACACAGAGA	1080
	ATGGAAGCCC	TGGAAAATCG	CCTGAGATAC	AGATGAAGAT	TAGAAATCGC	GACACATTTG	1140
25	TAGTCATTGT	ATCACGGATT	ACAATGAACG	CAGTGCAGAG	CCCCAAAGCT	CAGGCTATTG	1200
	TTAAATCAAT	AATGTTGTGA	AGTAAAACAA	TCAGTACTGA	GAAACCTGGT	TTGCCACAGA	1260
30	ACAAAGACAA	GAAGTATACA	CTAACTTGTA	TAAATTTATC	TAGGAAAAA	ATCCTTCAGA	1320
	ATTCTAAGAT	GAATTTACCA	GGTGAGAATG	AATAAGCTAT	GCAAGGTATT	TTGTAATATA	1380
	CTGTGGACAC	AACTTGCTTC	TGCCTCATCC	TGCCTTAGTG	TGCAATCTCA	TTTGACTATA	1440
35	CGATAAAGTT	TGCACAGTCT	TACTTCTGTA	GAACACTGGC	CATAGGAAAT	GCTGTTTTT	1500
	TGTAYTGGAC	TTTACCTTGA	TATATGTATA	TGGATGTATG	САТААААТСА	TAGGACATAT	1560
40	GTACTTGTGG	AACAAGTTGG	ATTTTTTATA	СААТАТТААА	ATTCACCACT	TCAGAGRAAA	1620
	АААААААА	AAAAAAAA	AAAAAAAA	АААААААА	AAANAAA		1668
45	(2) INFORM	ATION FOR SE	Q ID NO: 65	5:			
	(i)	SEQUENCE CH					
50		(B) TYP (C) STR	GTH: 1353 b E: nucleic ANDEDNESS: O DLOGY: line	acid double			
55	(xi)) SEQUENCE I	DESCRIPTION	: SEQ ID NO	: 65:		
	GGGTCGACCC	ACGCGTCCGC	CCACGCGTCC	GGATGGCTGC	GCTGTTGCTG	AGACACGTTG	60
	GTCGTCATTG	CCTCCGAGCC	CACTTTAGCC	CTCAGCTCTG	TATCAGAAAT	GCTGTTCCTT	120

TGGGAACCAC GGCCAAAGAA GAGATGGAGC GGTTCTGGAA TAAGAATATA GGTTCAAACC

	GTCCTCTGTC TCCCCACATT ACTATCTACA GTTGGTCTCT TCCCATGGCG ATGTCCATCT	240
_	GCCACCGTGG CACTGGTATT GCTTTGAGTG CAGGGGTCTC TCTTTTTGGC ATGTCGGCCC	300
5	TGTTACTCCC TGGGAACTTT GAGTCTTATT TGGAACTTGT GAAGTCCCTG TGTCTGGGGC	360
	CAGCACTGAT CCACACAGCT AAGTTTGCAC TTGTCTTCCC TCTCATGTAT CATACCTGGA	420
10	ATGGGATCCG ACACTTGATG TGGGACCTAG GAAAAGGCCT GAAGATTCCC CAGCTATACC	480
	AGTCTGGAGT GGTTGTCCTG GTTCTTACTG TGTTGTCCTC TATGGGGGCTG GCAGCCATGT	540
15	GAAGAAAGGA GGCTCCCAGC ATCATCTTCC TACACATTAT TACATTCACC CATCTTTCTG	600
13	TTTGTCATTC TTATCTCCAG CCTGGGAAAA GTTCTCCTTA TTTGTTTAGA TCCTTTTGTA	660
	TTTTCAGATC TCCTTGGAGC AGTAGAGTAC CTGGTAGACC ATAATAGTGG AAAAGGGTCT	720
20	AGTTTTCCCC TTGTTTCTAA AGATGAGGTG GCTGCAAAAA CTCCCCTTTT TTGCCCACAG	780
	CTTGCCTACT CTCGGCCTAG AAGCAGTTAT TCTCTCTCCA TATTGGGCTT TGATTTGTGC	840
25	TGAGGGTCAG CTTTTGGCTC CTTCTTCCTG AGACAGTGGA AACAATGCCA GCTCTGTGGC	900
25	TTCTGCCCTG GGGATGGGCC GGGTTGGGG GTGGGTTGGT GAGGCTTTGG GTGCCACTGC	960
	CTGTGGGTTG CTGGCTTAAA GGACAATTCT CTTCATTGGT GAGAGCCCAG GCCATTAACA	1020
30	CCTACACAGT GTTATTGAAA GAAGAGAGGT GGGGGTGGAG GGGAATTAGT CTGTCCCAGC	1080
	TAGAGGGAGA TAAAGAGGGC TAGTTAGTTC TTGGAGCAGC TGCTTTTGAG GAGAAAATAT	1140
35	ATAGCTTTGG ACACGAGGAA GATCTAGAAA ATTATCATTG AACATATTAA TGGTTATTTC	1200
33	TTTTTCTTGG ATTTCCAGAA AAGCCTCTTA ATTTTATGCT TTCTCATCGA AGTAATGTAC	1260
	CCTTTTTTTC TGAAACTGAA TTAAATACTC ATTTTATCTT TGAAAAAAAA AAAAAAAACC	1320
40	TNGGGGGGGG CCCCGGACCC NAATTGGCCC TAT	1353
45	(2) INFORMATION FOR SEQ ID NO: 66:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 1011 base pairs (B) TYPE: nucleic acid	
50	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 66:	
55	CGGAAGAAAG CAGCCATCCA GACATTTCAG AACACGTACC AGGTGTTAGC TGTGACCTTC	60
	AATGACACAA GTGATCAGAT TATTTCTGGT GGAATAGACA ATGATATCAA GGTCTGGGAC	120
	TGCGCCAGAA CAAGCTAACC TACACCATGA GAGGCCATGC AGATTCAGTG ACTGGCCTGA	180

	GTTTAAGTTC	TGAAGGCTCT	TATCTTTTGT	CCAATGCAAT	GGACAATACA	GTTCGTGTCT	240
	GGGATGTCCG	GCCATTTGCC	CCCAAAGAGA	GATGTGTAAA	GATATTTCAA	GGAAATGTGC	300
5	ACAACTTTGA	AAAGAACCTT	CTGAGATGTT	CTTGGTCACC	TGATGGAAGC	AAAATAGCAG	360
	CTGGCTCAGC	CGACAGGTTT	GTTTATGTGT	GGGATACCAC	AAGCAGGAGA	ATATTGTATA	420
10	AGCTGCCCGG	CCATGCTGGC	TCCATCAATG	AAGTGGCTTT	CCACCCTGAT	GAGCCCATCA	480
10	TTATCTCAGC	ATCGAGTGAC	AAGAGACTGT	ATATGGGAGA	GATTCAGTGA	AGATATGGAC	540
	TGGAAGACTC	CAAGGCCGCT	TGTCTTTGAG	ACCTCAGACT	GCATAAGTGA	TGCCAAATGT	600
15	TGGATGTCCA	GGYTAGCACC	CTCCCTTCAG	ATGACCATTG	CTAGCAAGAA	ACAGGAGGCG	660
	GTGGCCATAT	TCCAAAAACC	ACTTCTGTCC	CATTTCACCA	GGATGACTAA	GGCAAGCTCC	720
20	CTGTGGCCTC	TAAAAACCAC	CTGCCAGATT	TCAGGGACTG	TTTTTTTTT	TCTTTTTCTT	780
20	TTTTCCTGTT	TTCTAATGCA	GGCCCAATGT	GACAAATTTG	TTGGTTGGGA	TTTTTTTTT	840
	TTTTTGTAAC	TGGCTTGTAT	GATATTTTCT	TTCTGTATTT	CTCTATATCA	TTTTGTATTA	900
25	AAAGCCAAAT	AGATGCCTTT	TTACAAGARM	АААААААА	AAAAAAAAA	NNAAAAAAA	960
	CTGGGAGGG	GGGCCCGGTA	CCCAAATCGC	CGGATATGAT	CGTAAACAAT	С	1011
30							
<i>_</i>							
50	(2) INFORM	ATION FOR SE	EQ ID NO: 67	7 :			
		SEQUENCE CE (A) LENG (B) TYP (C) STR	~	ICS: ase pairs acid double			
35	(i)	SEQUENCE CE (A) LENG (B) TYP (C) STR	HARACTERIST: GTH: 1193 b E: nucleic ANDEDNESS: OLOGY: line	ICS: ase pairs acid double ar	: 67:		
35	(i)	SEQUENCE CI (A) LEN (B) TYP (C) STR (D) TOP	HARACTERIST: GTH: 1193 b E: nucleic ANDEDNESS: OLOGY: line DESCRIPTION	ICS: ase pairs acid double ar : SEQ ID NO		CCCGCGGGAC	60
35	(i) (xi GGCCGGGCGG	SEQUENCE CI (A) LEN (B) TYP (C) STR (D) TOP	HARACTERIST: GTH: 1193 b E: nucleic ANDEDNESS: OLOGY: line DESCRIPTION GGGCGCATCC	ICS: ase pairs acid double ar : SEQ ID NO	GCCGTCCGTG		60 120
35	(i) (xi GGCCGGGCGG CTGACAGCCG	SEQUENCE CI (A) LEN (B) TYP (C) STR (D) TOP (D) SEQUENCE I TGCGCACTGC GGTCAGAGGG	HARACTERIST: GTH: 1193 b E: nucleic ANDEDNESS: OLOGY: line DESCRIPTION GGGCGCATCC CGAACTGTGC	ICS: ase pairs acid double ar : SEQ ID NO CTGCCCCGGC	GCCGTCCGTG		120
35	(i) (xi GGCCGGGCGG CTGACAGCCG TGTCCCCAGA	SEQUENCE CI (A) LEN (B) TYP (C) STR (D) TOP (D) SEQUENCE I TGCGCACTGC GGTCAGAGGG	HARACTERIST: GTH: 1193 b E: nucleic ANDEDNESS: OLOGY: line DESCRIPTION GGGCGCATCC CGAACTGTGC GTCCTGGAAA	ICS: ase pairs acid double ar : SEQ ID NO CTGCCCCGGC TCAGGCCCGG	GCCGTCCGTG GCTGGACGCA AAAGGAACGG	GAGCCAGAGC AAGAAAGAGG	120
35 40 45	(xi GGCCGGGCGG CTGACAGCCG TGTCCCCAGA AGAGGCAGCG	SEQUENCE CI (A) LEN (B) TYP (C) STR (D) TOP) SEQUENCE I TGCGCACTGC GGTCAGAGGG GGAGCAGAGG	HARACTERIST: GTH: 1193 b E: nucleic ANDEDNESS: OLOGY: line DESCRIPTION GGGCGCATCC CGAACTGTGC GTCCTGGAAA GCAGGCCTTG	ICS: ase pairs acid double ar : SEQ ID NO CTGCCCCGGC TCAGGCCCGG GGAAGCTGAA TGGCCCAGCA	GCCGTCCGTG GCTGGACGCA AAAGGAACGG CCCGCCTGCC	GAGCCAGAGC AAGAAAGAGG AGGCGCTCGG	120 180
35 40 45	(xi GGCCGGGCGG CTGACAGCCG TGTCCCCAGA AGAGGCAGCG GGGCCGAACT	SEQUENCE CI (A) LEN (B) TYP (C) STR (D) TOP) SEQUENCE I TGCGCACTGC GGTCAGAGGG GGAGCAGAGG TCTGCGGGAG	HARACTERIST: GTH: 1193 b E: nucleic ANDEDNESS: OLOGY: line DESCRIPTION GGGCGCATCC CGAACTGTGC GTCCTGGAAA GCAGGCCTTG TACCTCTGCA	ICS: ase pairs acid double ar : SEQ ID NO CTGCCCCGGC TCAGGCCCGG GGAAGCTGAA TGGCCCAGCA GATGGGCCCA	GCCGTCCGTG GCTGGACGCA AAAGGAACGG CCCGCCTGCC AAAGCACAAG	GAGCCAGAGC AAGAAAGAGG AGGCGCTCGG AACTGGAGGT	120 180 240
35 40 45	(xi GGCCGGGCGG CTGACAGCCG TGTCCCCAGA AGAGGCAGCG GGGCCGAACT TTCAGAAGAC	SEQUENCE CI (A) LEN (B) TYP (C) STR (D) TOP (SEQUENCE I TGCGCACTGC GGTCAGAGGG GGAGCAGAGG TCTGCGGGAG GGCCTGGGAC GGCCTGGGAC GAGGCAGACG	HARACTERIST: GTH: 1193 b E: nucleic ANDEDNESS: OLOGY: line DESCRIPTION GGGCGCATCC CGAACTGTGC GTCCTGGAAA GCAGGCCTTG TACCTCTGCA TGGCTCCTGC	ICS: ase pairs acid double ar : SEQ ID NO CTGCCCCGGC TCAGGCCCGG GGAAGCTGAA TGGCCCAGCA GATGGGCCCA	GCCGTCCGTG GCTGGACGCA AAAGGAACGG CCCGCCTGCC AAAGCACAAG TGACAGTGAC	GAGCCAGAGC AAGAAAGAGG AGGCGCTCGG AACTGGAGGT	120 180 240 300 360
35	(xi GGCCGGGCGG CTGACAGCCG TGTCCCCAGA AGAGGCAGCG GGGCCGAACT TTCAGAAGAC ATGAGCACTT	SEQUENCE CI (A) LEN (B) TYP (C) STR (D) TOP (SEQUENCE I TGCGCACTGC GGTCAGAGGG GGAGCAGAGG TCTGCGGGAG GGCCTGGGAC GGCCTGGGAC GAGGCAGACG	HARACTERIST: GTH: 1193 b E: nucleic ANDEDNESS: OLOGY: line DESCRIPTION GGGCGCATCC CGAACTGTGC GTCCTGGAAA GCAGGCCTTG TACCTCTGCA TGGCTCCTGC CTGGCTACC	ICS: ase pairs acid double ar : SEQ ID NO CTGCCCCGGC TCAGGCCCGG GGAAGCTGAA TGGCCCAGCA GATGGGCCCA TGCACATGTA TGGAGGGGCCT	GCCGTCCGTG GCTGGACGCA AAAGGAACGG CCCGCCTGCC AAAGCACAAG TGACAGTGAC GCAGGGCCGG	GAGCCAGAGC AAGAAAGAGG AGGCGCTCGG AACTGGAGGT AAGGTTCCCG GCCCGAGAGC	120 180 240 300 360
35 40 45	(xi GGCCGGGCGG CTGACAGCCG TGTCCCCAGA AGAGGCAGCG GGGCCGAACT TTCAGAAGAC ATGAGCACTT TGACGGTGCA	SEQUENCE CI (A) LEN (B) TYP (C) STR (D) TOP) SEQUENCE I TGCGCACTGC GGTCAGAGGG GGAGCAGAGG TCTGCGGGAG GGCCTGGGAC GAGGCAGACG CTCCACCCTG	HARACTERIST: GTH: 1193 b E: nucleic ANDEDNESS: OLOGY: line DESCRIPTION GGGCGCATCC CGAACTGTGC GTCCTGGAAA GCAGGCCTTG TACCTCTGCA TGGCTCCTGC CTGGCCTACC GCCTGATGCG	ICS: ase pairs acid double ar : SEQ ID NO CTGCCCCGGC TCAGGCCCGG GGAAGCTGAA TGGCCCAGCA GATGGGCCCA TGCACATGTA TGGAGGGGCCT GGAGGGGGCT GGAGCTGGAT	GCCGTCCGTG GCTGGACGCA AAAGGAACGG CCCGCCTGCC AAAGCACAAG TGACAGTGAC GCAGGGCCGG GAGGAGGGCCT	GAGCCAGAGC AAGAAAGAGG AGGCGCTCGG AACTGGAGGT AAGGTTCCCG GCCCGAGAGC CTGATCCCCC	120 180 240 300 360 420

	TCCGGCGGTG GGGGCCGGGT TCACCAGCAG GGCAGCGGCT GAGCAAGGGC TTTCAGCTCC	660
5	TCCGGTGGTG GGGGCCGGGA TCACCAGCAC CAGAGCCTCG CAAGGGCCCC TTCCCTCCTC	720
5	CAGACCCTCC TTGGCCGGTG ACGCTGTGAC AGTGATGGCA GGTTCAGTGC CTTCAGCGCA	780
	GAGCGTGGAT GCTCTGGAAT CACCCGGACC CCTGGCCTTG GAGGGACCCT CCAGCCCCAG	840
10	GAATCTGCTT TGGAGGGAAA TGTCTATTTT TCTACCGGGA ATATTTTAGA GATTGGGGCA	900
	TGCTGGCTCC TCCCGCCAGC TGCAAACCTG CACCTTCCGC CTGATTCCCG ATCCCCCTGC	960
15	GTGGGCCGCA TTCCTGGTCC CCTGCCTGCG TCCATCGAGG GGCCTGGCTG TGGCCTGTTT	1020
13	TCCTTTGACC CCACACAGCG TCATTGCGGG TCATGGGGAG CCCCTGGTGG GAGCTTGTGG	1080
	AGTCGGATCA CGTACCTGTG CAGAAACCGC CTCTGTGGCT GCATTTGAAA TAAAACCCGA	1140
20	CCCAGCAGCA AAAAAAAAA AAAAAANCNC NAGGGGGGGC CCGGNACCCA ATT	1193
25	(2) INFORMATION FOR SEQ ID NO: 68:	
	(i) SEOUENCE CHARACTERISTICS:	
	(A) LENGTH: 560 base pairs	
30	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 68:	
35	GAATTCGGCA CGAGTTGGCA CATGATGCAA AATGCATTTC TCAGAGTAGA TTGCAGTCAA	~ ^
		60
	AAATGTTGGA AACTACTAAG CATGTGCARA TAGCATGCAT GCTGCTGCTG ACCTGCCAGA	120
40	AAATGTTGGA AACTACTAAG CATGTGCARA TAGCATGCAT GCTGCTGCTG ACCTGCCAGA TATTTCTCCC TTCCTCCCTT TCTCCCTCAT TTATTCATTC	
40		120
40	TATTTCTCCC TTCCTCCCTT TCTCCCTCAT TTATTCATTC	120 180
40 45	TATTTCTCCC TTCCTCCCTT TCTCCCTCAT TTATTCATTC	120 180 240
	TATTTCTCCC TTCCTCCCTT TCTCCCTCAT TTATTCATTC	120 180 240 300
45	TATTTCTCCC TTCCTCCCTT TCTCCCTCAT TTATTCATTC	120 180 240 300 360
	TATTTCTCCC TTCCTCCCTT TCTCCCTCAT TTATTCATTC	120 180 240 300 360 420
45	TATTTCTCCC TTCCTCCCTT TCTCCCTCAT TTATTCATTC	120 180 240 300 360 420 480

(2) INFORMATION FOR SEQ ID NO: 69:

(i) SEQUENCE CHARACTERISTICS:

220

(A) LENGTH: 1657 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear

5

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 69: CGGACNGAGC CGCCGCCGGG CACTTCCTGT GGAGGCCGCA GCGGGTGCGG GCGCCGACGG 60 10 GCGAGAGCCA GCGAGCGAGC GAGCGAGCCG AGCCGAGCCT CCCGCCGTCG CCATGGGCCA 120 GAACGACCTG ATGGGCACGG CCGAGGACTT CGCCGACCAG TTCCTCCGTG TCACAAAGCA 180 GTACCTGCCC CACGTGGCGC GCCTCTGTCT GATCAGCACC TTCCTGGAGG ACGGCATCCG 240 15 TATGTGGTTC CAGTGGAGCG AGCAGCGCGA CTACATCGAC ACCACCTGGA ACTGCGGCTA 300 CCTGCTGGCC TCGTCCTTCG TCTTCCTCAA CTTGCTGGGA CANTGACTGG CTGCGTCCTG 360 20 GTGTTGAGCA GGAACTTCGT GCAGTACGCC TGCTTCGGGC TCTTTGGAAT CATAGCTCTG 420 CAGACGATTG CCTACAGCAT TTTATGGGAC TTGAAGTTTT TGATGAGGAA CCTGGCCCTG 480 GGAGGAGGCC TGTTGCTGCT CCTAGCAGAA TCCCGTTCTG AAGGGAAGAG CATGTTTGCG 540 25 GGCGTCCCCA CCATGCGTGA GAGCTCCCCC AAACAGTACA TGCAGCTCGG AGGCAGGGTC 600 TTGCTGGTTC TGATGTTCAT GACCCTCCTT CACTTTGACG CCAGCTTCTT TTCTATTGTC 660 30 CAGAACATCG TGGGGCACAG CTCTGATGAT TTTAGTGGCC ATTGGTTTTA AAACCAAGCT 720 GGCTGCTTIG ACTCTTGTTG TGTGGCTCTT TGCCATCAAC GTATATTTCA ACGCCTTCTG 780 GACCATTCCA GTCTACAAGC CCATGCATGA CTTCCTGAAA TACGACTTCT TCCAGACCAT 840 35 GTCGGTGATT GGGGGCTTGC TCCTGGTGGT GGCCCTGGGC CCTGGGGGTG TCTCCATGGA 900 TGAGAAGAAG AAGGAGTGGT AACAGTCACA GATCCCTACC TGCCTGGCTA AGACCCGTGG 960 40 CCGTCAAGGA CTGGTTCGGG GTGGATTCAA CAAAACTGCC AGCTTTTATG TATCCTCTTC 1020 CCTTCCCCTC CCTTGGTAAA GGCACAGATG TTTTGAGAAC TTTATTTGCA GAGACACCTG 1080 AGAATCAATG GCTTCAGGAC ATGGGTTCTC TTCTCCTGTG ATCATTCAAG TGCTCACTGC 1140 45 ATGAAGACTG GCTTGTCTCA GTGTTTCAAC CTCACCAGGG CTGTCTCTTG GTCCACACCT 1200 CGCTCCCTGT TAGTGCCGTA TGACAGCCCC CATCAAATGA CCTTGGCCAA GTCACGGTTT 1260 50 CTCTGTGGTC AAGGTTGGTT GGCTGATTGG TGGAAAGTAG GGTGGACCAA AGGAGGCCAC 1320 GTGAGCAGTC AGCACCAGTT CTGCACCAGC AGCGCCTCCG TCCTAGTGGG TGTTCCTGTT 1380 1440 55 GATAAGTGGG ATCTACCAAT TGATTCTGGC AAAACAATTT CTAAGATTTT TTTGCTTTAT 1500 GTGGGAAACA GATCTAAATC TCATTTTATG CTGTATTTTA TATCTTAGTT GTGTTTGAAA 1560 60 1620

	AAAAAAACTC GRGGGGGGC CCGGTACCCA AATCGCC	1657
5		
	(2) INFORMATION FOR SEQ ID NO: 70:	
10	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 711 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 70:	
	GGCACGAGCG AAGACCCTGT TCGGACCCTG CCCCGATTCC AGACTCAGGT AGATCGTCGG	60
	CATACCCTCT ACCGTGGACA CCAGGCAGCC CTGGGGCTGA TGGAGAGAGA TCAGGTATCC	120
20	CCCAGGGAGT AGGGGCTACC TTGAGGGGAT GATAGACCTC CCCCACTCCC AGTGKKACTC	180
	TGGAAATATG AAGGAACTAG GGAGTGGAAG AGATTTCAGA GCTGGGGAGA GGAGTTCCTC	240
25	CCTTCAAAGC CAGCAACTGC CTTTGGGGAA TGTCGGGGGG TCTCTCCTTT CTCCTGCTTG	300
	TTTRAGGTGG TACACAGTCC CCCCTTCAMC TGGSGGGAAG CTGTNCCGGA CARACTCATC	360
20	TCAGCTTTCC CTTGGGGCAG GATCGGGGGC AGCAGCTCCA GCAGAAACAG CAGGATCTGG	420
30	AGCAGGAAGG CCTCGAGGCC ACACAGGGGC TGCTGGCCGG CGAGTGGGCC CCACCCCTCT	480
	GGRAGCTGGG CAGCCTCTTC CAGGCCTTCG TGAAGAGGGA GAGCCAGGCT TATGCGTAAG	540
35	CTTCATAGCT TCTGCTGGCC TGGGGTGGAC CCAGGACCCC TGGGGCCTGG GTGCCCTGAG	600
	TGGTGGTAAA GTGGAGCAAT CCCTTCACGC TCCTTGGCCA TGTTCTGAGC GGCCAGCTTG	660
40	GCCTTTGCCT TAATAAATGT GCTTTATTTT CAAAAAAAA AAAAAAAAAC T	711
45	(2) INFORMATION FOR SEQ ID NO: 71: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 935 base pairs (B) TYPE: nucleic acid	
50	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
-	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 71:	
	GGCACAGGGT GAAAGCCAGC TAAACCCCAA GTGGAGAAGT GAAAGACATG GTTGTTCCCA	60
55	TAAGTTTATT GCTCACATTA TGAAAGAAGC CATAGTCATG AGTGAACCAC TCCCTAGGTT	120
	GATAAGGAAA CCAACACGGA AGATCTCTTT CTGGAAGAAG CAGCCAGCCT CGTGAAGGAG	180
60	CGGCCCAGCC GCCGGGCCCG AGGGTCGCCT TTTGTTCGGA GTGGCACGAT TGTCCGTTCC	240

	CAGACATTCT	CGCCTGGAGC	ACGAAGCCAG	TATGTTTGCA	GACTTTATCG	TAGTGACAGC	300
5	GACAGTTCAA	CGCTGCCCCG	GAAGTCCCCC	TTTGTCCGAA	ATACTTTGGA	AAGACGAACC	360
	CTTCGCTATA	AGCAGTCATG	CAGGTCTTCC	CTGGCTGAGC	TCATGGCCCG	CACCTCCCTG	420
	GACTTGGAGC	TGGATCTCCA	GGCGTCGAGA	ACACGGCAGA	GGCAGCTGAA	TGAGGAGCTC	480
10	TGCGCCCTCC	GTGAGCTGCG	GCAGCGGTTN	GGAGGACGCC	CAGCTCCGTG	GCCAGACTGA	540
	CCTCCCACCC	TGGGTGCTTC	GGGACGAGCG	GCTCCGTGGC	CTGCTGCGGG	AGCCGAGCGG	600
15	CAGACAAGAC	AGACCAAACT	TGACTACCGT	CATGAGCAGG	CGGCTGAGAA	GATGCTGAAG	660
•	AAGGCCTCCA	AGGAGATCTA	CCAGCTGCGT	GGCAGAGCCA	CAAAGAGCCC	ATCCAAGTGC	720
	AGACCTTTAG	GGAGAAGATA	GCATTCTTCA	CAAGGCCAAG	GATCAACATA	CCTCCTCTCC	780
20	CAGCCGACGA	CGTCTGATGG	AGTGCATTGT	GCACATGAAG	TATTTATCCA	CCTGTTTTAT	840
	TTTCATGAAG	TTCTTAGACT	AGCTGAATTT	GTCTTTAAAA	TATTTGTGCA	AAGCTATTAA	900
25	TATACACATT	TTGTAAAAA	АААААААА	AAACT			935
	(2) INFORM	ATION FOR SE	EO ID NO: 72	2:			
30		SEQUENCE CI	_				
		(A) LEN	GTH: 504 ba E: nucleic	se pairs			
35			ANDEDNESS: OLOGY: line				
	(xi) SEQUENCE I	DESCRIPTION	: SEQ ID NO	: 72:		
						GCTCTGACTC	60
40						GCACGGACTG	120
					CATTGAGCCC		180
45					GTTCTTCAGC		240
						CACTITCCGT	300
						GATGAAGACT	360
50						AGGATTATCG	420
						GARAAAAAA	480
55		GGGGGGGGC			COLC MUNTI	G. HARMANA	504
-	,=== =						304

(2) INFORMATION FOR SEQ ID NO: 73:

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5	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 620 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 73:	
10	GAATTCGGCA CGAGGAGGAG GGGAGGCGGG GTAAGTTTGG TGGGAAACTC TGTAATTTCC	60
	WTTTTACTT TCACAGCAAT AGTGCAGAAT CCAGAATGGA TGTCCTCTTT GTAGCCATCT	120
15	TTGCTGTGCC ACTTATCCTG GGACAAGAAT ATGAGGATGA AGAAAGACTG GGAGAGGATG	180
15	AATATTATCA GGTGGTCTAT TATTATACAG TCACCCCCAG TTATGATGAC TTTAGTGCAG	240
	ATTTCACCAT TGATTACTCC ATATTTGAGT CAGAGGACAG GCTGAACAGG TTGGATAAGG	300
20	ACATAACAGA AGCAATAGAG ACTACCATTA GTCTTGAAAC AGCACGTGCA GACCATCCGA	360
	AGCCTGTAAC TGTGAAACCA GTAACAACGG AACCTCAGAG TCCAGATCTG AACGATGCCG	420
	TGTCCAGTTT GCGAAGTCCT ATTCCCCTCC TCCTGTCGTG TGCCTTTGTT CAGGTGGGA	480
25	TGTATTTCAT GTAGAAGGTG GAAGAAGGCT GCTATGACTC TTTGGATGGG AGTCTGGCAA	540
	GAGGAAATTG GAAGATAAAA TAAATAATAA GTGAAATAAA AAAAAAAA	600
30	GGGGGGGCCC GGTACCCAAT	620
35	(2) INFORMATION FOR SEQ ID NO: 74: (i) SEQUENCE CHARACTERISTICS:	
40	(A) LENGTH: 581 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 74:	
45	ACAAGGTGTG TGTAAAGTTT ATGTTTGTAA ACTGAATTCT ATCTTAAATC CAAAAAGAAC	60
	TCGGGAGTAA TTCATTTTTG TAGCATAAAG ATCCCTAAGT TTTATTTTGA AATATCTGAT	120
50	TTTTACACGT TAAAAAATAA CAGGGCATCG AGAGGATTCC TAGGTGACAT CCAGACTCCT	180
	TTAGCTTTGT GTGTGTGGCA CCGGTTAGTC TGCTTCTCT TCCTTTCTTG CACTGCTTCA	240
	CACAGCCATG CCCTGCCAGC CCGGGCAGGT GCCTTCCTGT CAATGTACAT TTGGGCTTCT	300
55	GCTCATGCTG CCCTCCCTCC CCTCCCCTGC CTCCCAACCC CGCCCCTTTT GTTCCTCCAT	360
	GGAGTACTTC CATGGGTGTG CCTCCCCCAG CCAAGCCATA ATAGGTGGTT TCCCCTTCGC	420
	TTCTGTAGCC CTTGCAGACA TCCTCTGTTT ACAGTAGGTG TTGACTTACT TCCCCTCTCC	480

	CCGSTAAAGC CATAAACTCC TTAAGGACAG GTAGCATTCT TAGTATCTTC GTTCTTCTCA	540
	ATGACCAGTA GACCATTAAA CATGTAGCAA ACAAATGTGA A	581
5		
	(2) INFORMATION FOR SEQ ID NO: 75:	
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 1843 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 75:	
	AAACCCAACN CCCTCCGGTC CCCNAAAGAA AGCCCAGCCC AAATCCCAAG CCGGCAGTGA	60
20	GCCCGCGAAC AAGGCCCTCA AGACGCCCAG NCGAACAAGC AGCCCCCAGG AGGCCCCGCA	120
	AGAGAACTCC CTGGCGGCCC AAGCGGGCAG CTTCTGTGCG GCAGAACTCA GCCACCGAGA	180
25	GCGCAGACAG CATCGAGATT TATGTCCCGG AGNCCCAGAC CAGGCTCTGA GACCATGCAG	240
	GAGGAAAGAA ACGATTTTAA ATCATTAAAA ACACAAAAAC TAAGTGCGAA CGGAACAGAG	300
	TTTTCTCAAC CTTTGCTATG GTTATTCTGT CTAGAGACCC TGAGCCAACT TTCAAATTGA	360
30	CGCATACAAG GGCTCACAAT TTGGCTTTTT TGGGTCCCTC CCAGCTTTAG GTTATGAAGA	420
	TTTTACTCAC AAAAAAAATC AACAAAA; ~ C ACGAAACTAG AAAACTTTTT TTTTCCTCTT	480
35	GCTGGCCGTG GTGGACTAGA TAGATGGACG TCGGCAACTC CCGGCCCAGC CTCCATACTG	540
	CGGTCTTTTT ACTCGTTCTA TCTGATGAGA ACTCACACTA GCTTGTTTAC AAGATGACGA	600
	CAGTCCAAGG GCAGCCTTGG GCACCTGCCA TGTCCCTCCT TTCCCCAGCT ATCCCCGCTC	660
40	TGACCTTGAT TTTCATTCTT ATGTTTTTCT CTTTTCCCTT CAGAGCTCAC ACAGTGGTCA	720
	CCATTGTGGC AAGCGGCTTT CTGGGTCTCA GCCCTCTCTG CGGTTGAGGG CCCAGAGGAC	780
45	AGAGAGATGG ACATGCGTCC CCTCCCTCCC CCCGCCAAGT GCTCACACAC AACCTCACGC	840
	GCACACACA ACACGCAGAT GGAGGCGCCT CACTGGGAGG TGCCCCGCCA GCCCTGGGCA	900
	GTGTCAGGCA GGACTCACTC ACCGCTGAGC AGATGAGAGA AGTTTTAGTC TTGGCGGGTG	960
50	GAAATGAGAC GAAGCCACAG TTATCACACT CCAGACTCCT GCCCTTTTAT TTTCTCCAGC	1020
	CCCTTCTTCC TTCAGCAAAA TCTAGGACTC CCGAGTGGCT TCCAGGGGGC CGTCAGTCCT	1080
55	CAGCCGCGCC TGTGTCCGGT GCCCGAGGGG CGGGCGGCGG TGTCTGTATG TATGTGTACA	1140
	TATGCACATA GACCTTAGAG TGTATAGTTA ACAAACGCCC ATCTGCTCAC CCATGCCCAC	1200
	CCAGCGCCGC CGCCGCTGGC TCTCGGGGGCA CCTGGCAGGA GGCGGGTGTG TGAATAGCAT	1260
60	ATATOTATION AND TO THE TOTAL TOTAL AND THE T	1320

	GTGTGTAAGC AGCCCTTTTT TTTTTTGGTC TCCACCCCCC TCCCCCCGCC CCGCACTCCT	1380
_	AAGGGCCCAT CTGCCCAGCC TCTGAGTTTT CTGTTCTATT TTTTTTTTAA CCCCAATTAT	1440
5	CCTTCTCTCT CTCCTGCCCC CGCATCCCAC TCCCAGGGTG TCACGAGCCC TGAGCTGCAA	1500
	TGGCCCGGGC CTGCAGGGCG GGGTAGGGGA GGGCARGGCT SAGCCCCGAA GCCAGCTCAG	1560
10	TACCTGAGGG GCTGCTCTAT GCTGTGTATG CGCCTCTCTG GCATCCGAGA CATCCTCTTG	1620
	GTGGCGCTTG CTNGCAGGGG ACCCCCCCC CGTCCCCAGG TGAACCAAGG GTCTGCTCCG	1680
15	GGGCCCATTT CCAGCTTGGC CGCCGTCTGT GACCTTGGGC AAGTCACTTG ACCTCTGTGT	1740
13	GCCTCAACTT CCTCCTCTGT AAAACGGGGA CAGTCCCTGC CCCTCCCTAC CTCACAGGCA	1800
	TGTTGTGAGA ATAAATGAGG TAACGTGTAA AAAAAAAAAA	1843
20		
	(2) INFORMATION FOR SEQ ID NO: 76:	
~ -		
25	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 1441 base pairs	
	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
20	(D) TOPOLOGY: linear	
30	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 76:	
	TCGACCCACG CGTCCGGCTC CCCGAGCCCT GCCAACCATG GTGAACTTGG GTCTGTCCCG	60
35	GGTGGACGAC GCCGTGGCTG CCAAGCACCC GGGACTCGGG GAGTATGCCG CATGCCAGTC	120
	ACACGCCTTC ATGAAGGGCG TTTTCACCTT CGTCACAGGC ACCGGCATGG CCTTTGGCTT	180
40	GCAGATGTTC ATTCAGAGGA AGTTTCCATA CCCTTTGCAG TGGAGCCTCC TAGTGGCCGT	240
40	GGTTGCAGGC TCTGTGGTCA GCTACGGGGT GACGAGAGTG GAGTCGGAGA AATGCAACAA	300
	CCTCTGGCTC TTCCTGGAGA CCGGGCAGCT CCCCAAAGAC AGGAGCACAG ATCAGAGAAG	360
	CCLCLOCCLC LLCCLGCACA CCGGGCAGCL CCCCAAAGAC AGGAGCACAG ALCAGAGAAA	300
45	CTAGGAGAGC TCCAGCAGGG GCACAGAGGA TTGGGGGCAG GAGGAGTCTG GAACACAGCC	420
45		
	CTAGGAGAGC TCCAGCAGGG GCACAGAGGA TTGGGGGCAG GAGGAGTCTG GAACACAGCC	420
4550	CTAGGAGAGC TCCAGCAGGG GCACAGAGGA TTGGGGGCAG GAGGAGTCTG GAACACAGCC TTCATGCCCC CTGACCCCAG GCCGACCCTC CCCACACCCT AGGGTACCCC AGTCGTATCC	420 480
	CTAGGAGAGC TCCAGCAGGG GCACAGAGGA TTGGGGGCAG GAGGAGTCTG GAACACAGCC TTCATGCCCC CTGACCCCAG GCCGACCCTC CCCACACCCT AGGGTACCCC AGTCGTATCC TCTGTCCGCA TGTKTGGCCA GGCCTGACAA ACACCTGCAG ATGGCTGCTG CCCCAACCTG	420 480 540
	CTAGGAGAGC TCCAGCAGGG GCACAGAGGA TTGGGGGCAG GAGGAGTCTG GAACACAGCC TTCATGCCCC CTGACCCCAG GCCGACCCTC CCCACACCCT AGGGTACCCC AGTCGTATCC TCTGTCCGCA TGTKTGGCCA GGCCTGACAA ACACCTGCAG ATGGCTGCTG CCCCAACCTG GGACCTGCCC AGRAGGTTGG AGCAGAAAGG GCTCTCCCTG GGGTGGTGTT TCTCCTCTAG	420 480 540

TTCTCTCCAC CCAAGTTTCC ACCTGACCAG GTGAAAAACA AATCAGAAGG GTAAGATGAT

	GACAGGTCAC ATGAAACCTT TATTACCCTA CAGTTGATAT ATGAGGATCA CATGCAAGTT	900
	ACATACTGAG GATGTACAGG GAAGTTCCCA GCGCTGAACC CCAGAATTAG ACGTTCGCAT	960
5	CAGCCCCGTA GGCCACGTGG ACACCACCAC AGCCTCTCTG TATGGGGGTC TGCCTCTGTA	1020
	GCACTTGGCA TGTAGGGGCA GAGCAAAAGG GGCCANGCTG GCCAGAGCCT GGCTGCTGGG	1080
10	NAGARGAGGG ACTTGTGGGS CACGCCACNT GCCTATCATT CCCCAYTCAT CTATTAGCCA	1140
	AAGTCACTCC CCAGAGGCAG AGCTAGCCCG TTGTAGCCGT GTCTGTGTGG AGGGAAAGCT	1200
	TCTGAGTGGG CAAGCCTACA CACAGCCCCG AGCCCCAAGA GGAGGAAGAG GTGGAGACCA	1260
15	GACGGAACCT CCACAAGTCC ATCATGGTTA CAGCTGGCTT CCCCGCAGCA CCGAAGACCC	1320
	ACAGCATNGG CCCTGCTGCC CCCGACCCAG CTCAGCTGCC ANGCCTCACC TTGCCAGGAA	1380
20	TTGAAAGAAA GTTATTGAGT ACTAATTGGC CTCAGAGTNA CAGGAAGCTC AAGTTAAAGT	1440
20	G	1441
25	(2) INFORMATION FOR SEQ ID NO: 77:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 910 base pairs	-
30	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
	(D) TOPOLOGY: linear	
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 77:	
	GGCAGAGCTG GCCTTCGACT CGCTATGTCC ACTAACAATA TGTCGGACCC ACGGAGGCCG	60
	AACAAAGTGC TGAGGTGAGG ACCCCAGCGT CGTGGGCACG GGTTCGGGTT GTGGGTGTGG	120
40	ATCGGGGCCC TGGGAAGCGC CTGTCTATCC CGGGGGCAGG ACCTGAGCGC CCCTGACCCT	180
	CGAGCCTGTC GCAGGTACAA GCCCCCGCCG AGCGAATGTA ACCCGGCCTT GGACGACCCG	240
45	ACGCCGGACT ACATGAACCT GCTGGGCATG ATCTTCAGCA TGTGCGGCCT CATGCTTAAG	300
	CTGAAGTGGT GTGCTTGGGT CGCTGTCTAC TGCTCCTTCA TCAGCTTTGC CAACTCTCGG	360
	AGCTCGGAGG ACACGAAGCA AATGATGAGT AGCTTCATGT GAGACTTGCC CTACAGAACA	420
50	AGTGACTCTT GAGTAAGGGG TGGGGGGACC CCAGCCTGGC CATCCTAGAC TGACACCTCT	480
	CTCCTGTCTT CATGCTGTCC ATCTCTGCCG TGGTGATGTC CTATCTGCAG AATCCTCAGC	540
55	CCATGACGCC CCCATGGTGA TACCAGCCTA GAAGGGTCAC ATTTTGGACC CTGTCTATCC	600
<i>55</i>	ACTAGGCCTG GGCTTTGGCT GCTAAACCTG CTGCCTTCAG CTGCCATCCT GGACTTCCCT	660
	GAATGAGGCC GTCTCGGTGC CCCCAGCTGG ATAGAGGGAA CCTGGCCCTT TCCTAGGGAA	720
60	CACCCTAGGC TTACCCCTCC TGCCTCCCTT CCCCTGCCTG CTGCTGGGGG AGATGCTCTC	780

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	CATGTTTCTA GGGGTATTCA TTTGCTTTCT CGTTGAAACC TGTTGTTAAT AAAGTTTTTC	840
_	ACTCTGAAAA AAAAAAAAA AAAAAAAAAC TYGRGGGGG GCCCGGAACC CAATTCSCCG	900
5	GATAGTGAGT	910
10		
	(2) INFORMATION FOR SEQ ID NO: 78:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 2776 base pairs	
15	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
	(D) TOPOLOGY: linear	
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 78:	
20	TCGACCCACG CGTCCGGGCG GGCAGTGATG GCGGCTGGTG ATGGGGACGT GAAGCTAGGC	60
	ACCCTGGGGA GTGGCAGCGA GAGCAGCAAC GACGGCGGCA GCGAGAGTCC AGGCGACGCG	120
25	GGAGCGGCAG CGRAAGGGGG AGGCTGGGCG GCGGCGGCGT TGGCGCTTCT GACGGGGGC	180
	GGGGAAATGC TGCTGAACGT GGCGCTGGTG GCTCTGGTGC TGCTGGGGGC CTACCGGCTG	240
20	TGGGTGCGCT GGGGGGCG GGGTCTGGGG GCCGGGGCCG GGGCGGCGA GGAGAGCCCC	300
30	GCCACCTCTC TGCCTCGCAT GAAGAAGCGG GACTTCAGCT TGGAGCAGCT GCGCCAGTAC	360
	GACGGCTCCC GCAACCCGCG CATCCTGCTC GCGGTCAATG GGAAAGTCTT CGACGTGACC	420
35	AAAGGCAGCA AGTTCTACGG CCCGGCGGGT CCATATGGAA TATTTGCTGG TAGGGATGCC	480
	TCCAGAGGAC TGGCCACATT TTGCCTAGAT AAAGATGCAC TTAGAGATGA ATATGATGAT	540
40	CTCTCAGATT TGAATGCAGT ACAAATGGAG AGTGTTCGAG AATGGGAAAT GCAGTTTAAA	600
40	GAAAAATATG ATTATGTAGG CAGACTCCTA AAACCAGGAG AAGAACCATC AGAATATACA	660
	GATGAAGAAG ATACCAAGGA TCACAATAAA CAGGATTGAA CTTTGTAAAC AACCAAAGTC	720
45	AGGGGCCTTC AGAACTGCAA TTCTTACTCC CTTTCACAGA CTGTCCGGAG TCTTTGGGTT	780
	TGATTCACCT GCTGCGAAAA ACATTCAACA AATTGTGTAC AAGATAAATT AATCTCACTA	840
50	TGAAGATTTG AATAACTAGA CATTATTTAT GCTGCCAAAC TCATTTGTTG CAGTTGTTTG	900
50	TAATGTCTAG TGGGGCTTCA TCATCCTGAA AAGAAGGAGA CAGGGATTTT TTTAAAGAGC	960
	AAGAAAGTCA CAATATTACT TCTTTCCTTC CTTTTTTCCT TCTTTCCTTT CTTCTT	1020
55	TTTCTTTCTT TTTAAAATAT ATTGAAGACA ACCAGATATG TATTTGCTAC TCAAGTGTAC	1080
	AGATCTCCTC AAGAAACATC AAGGGACTCC TGTGTCACAT ACTGTGTTTT TATTTTAACA	1140
60	TGGGTGAGGG AGGCGACCTG ATCAGGGGAG GTGGGGGTAC ACATCAATTT GAGTTGTTCA	1200
60		

	GGCTACTGAA	ACATTAAAAT	GTGAATTCCC	AAACTTTTCT	TTTTGGCTTT	GTCAGGGAAA	1260
	AGAAAAATAT	CTTTATAAAG	AAATCTTTGG	AAATTAGGAG	AAGGAATTTC	AGGTGGGTTT	1320
5	AAGTCAGAGC	TAGTTCCCCA	ACAGAAAGAT	CATTTGAAAC	CAGTTTTTAT	CCCTTCTCTT	1380
	TCCTTCCCTT	TCCCTAAATC	AAATCAATAT	TAATTGTGCC	TTATTTCACT	TAACATAGAC	1440
10	TTGAATTATT	TTTAGGGAAA	GCCCCTATAA	TGAATTCAGA	AATCACTACA	AGCAGCATTA	1500
	AGACTGAAGT	TGGAATATTC	TGTTGACCAT	AAAACCTTGA	TATCATTCTG	TGTATATAGA	1560
	ATGTAAAAGG	AATATTACAG	TGTTAACTGC	CATATATGTA	ATATACACAA	ACTCAATTAG	1620
15	CATTGTAATG	GCCAAATGCA	TTCCCCCATG	CTTTTCTGTT	TTCAAAAAAA	TTGAAAAACA	1680
	AATCAACTCT	TATCCCCAAC	AGCTGCCTAA	TTTTAGGAGT	CTGACCCTCC	ACATCTCACT	1740
20	GGTGTGGGTG	CATGGGGCTG	TGGAGTGGGT	GTCAGTATGG	ATGTGTCTGA	ATGTGTGAGG	1800
	CCTTGGAAGG	GACTCTTTCT	GCAGATACTG	TAAATACAAG	TACCATTTTA	ATAAAGCATG	1860
	TACAATAAAC	CAAAATAAGC	TTGAGTTGGA	CTTTATATAC	AGAACTGTAA	GCCAGTGCAT	1920
25	TATGATACAG	TTGTAAGATT	GTGCATTTGA	TTCAAGATAA	GGAAAAATCT	TGGAAATGAA	1980
	AAGCAGGCAC	KGGTTAACCA	AGTTGTACAC	ATTGTACCAC	ATTCAGCATA	ACTTTAGGAA	2040
80	GAAATTCCAC	TTTGTGAACA	TTCTCCAGAA	ATCCAAGATT	ATTCAGGTAA	GAATTGGTAT	2100
	ATTAAATGTA	CATCTTTTTA	CTTTCTATTT	TGATGCCAAC	TGATTATACT	AGACAATTAG	2160
	CACTCCAGGT	GGTTATTGAA	CACAAAACAG	TAAAAGAATA	TTGCACTGAT	AGATACTAAA	2220
35	TTATTATTTT	ATTAGGTTGA	AAAAGCCCTT	ACTAAAAGCC	CCTCATATAT	CAATTACTTT	2280
	ATTTCATTAT	GACTACTTAG	GTTCCGGGCT	GGGGACAAGT	TCACTTAAAA	AGGCAATGTT	2340
Ю	ATTTAACAGG	TCACCAGTTA	AGACTTCTGC	TTTGTAGATA	CATGCAGAAG	CCATCAAACA	2400
	AGGGGGRGCT	TTTAACTGCA	ACAATAAGCT	AAAGTATGTA	AAATACTACA	TTCTATTCAG	2460
	TCTTGGAGTG	TTTTGTAGAA	AGTTATCTTC	AGCCAAATCT	TTGCTGAAGA	CTGGTTGTGG	2520
15	AGTGTTGGTA	AATGCTTTGT	GTTTTTATGT	AAAATATTTT	CTAAACAAAA	AATGTTAAAA	2580
	GTACATGTCC	TCTGTAGTAA	ACTGATATCT	ATATATATGA	ATCATTCAAG	CCTAAAGTCT	2640
50	AGTAATAAAC	TGTACTTGTG	AATAGAGAAA	CCCTAAATAT	TCATGCAGWA	AAAATTATGC	2700
	GGTCTGTTAA	GAAAAATGAG	TAATTTGTGT	TTTGGACTTG	AAATAAACAG	TGTTCTGTAG	2760
	ATAATTCCTC	AACTTC					2776

(2) INFORMATION FOR SEQ ID NO: 79:

60 (i) SEQUENCE CHARACTERISTICS:

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(A) LENGTH: 1525 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 79:

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	(XI) SEQUENCE DESCRIPTION. SEQ ID NO. 77.	
	CCGCTGCTGA TAACTATGGC ATCCCCCGGG CCTGCAGGAA TTCGGCACGG AGCTACGGCG	60
10	CCGCCTGGCT CCTGCTGNCA CCTGCAGGCT CGTCGCGGGT GGAGCCCACC CAAGACATCA	120
	GCATCAGCGA CCAGCTGGGG GGCCAGGACG TGCCCGTGTT CCGGAACCTG TCCCTGCTGG	180
15	TGGTGGGTGT CGGCGCCGTG TTCTCACTGC TATTCCACCT GGGCACCCGG GAGAGGCGCC	240
13	GGCCGCATGC GGASGAGCCA GGCGAGCACA CCCCCCTGTT GGCCCCTGCC ACGGCCCAGC	300
	CCCTGCTGCT CTGGAAGCAC TGGCTCCGGG AGCSGGCTTT CTACCAGGTG GGCATACTGT	360
20	ACATGACCAC CAGGCTCATC GTGAACCTGT CCCAGACCTA CATGGCCATG TACCTCACCT	420
	ACTCGCTCCA CCTGCCCAAG AAGTTCATCG CGACCATTCC CCTGGTGATG TACCTCAGCG	480
25	GCTTCTTGTC CTCCTTCCTC ATGAAGCCCA TCAACAAGTG CATTGGGAGG AACATGACCT	540
23	ACTICTCAGG CCTCCTGGTG ATCCTGGCCT TTGCCGCCTG GGTGGCGCTG GCGGAGGGAC	600
	TGGGTGTGGC CGTGTACGCA GCGGCTGTGC TGCTGGGTGC TGGCTGTGCC ACCATCCTCG	660
30	TCACCTCGCT GGCCATGACG GCCGACCTCA TCGGTCCCCA CACGAACAGC GGACTKTCGT	720
	GTACGGCTCC ATGAGCTTCT TGGATAAGGT GGCCAATGGG CTGGCAGTCA TGGCCATCCA	780
35	GAGCCTGCAC CCTTGCCCCT CAGAGCTCTG CTGCAGGGCC TGCGTGAGCT TTTACCACTG	840
23	GGCGATGGTG GCTGTGACGG GCGGCGTGGG CGTGGCCGCT GCCCTGTGTC TCTGTAGCCT	900
	CCTGCTGTGG CCGACCCGCC TGCGACGCTG GGACCGTGAT GCCCGGCCCT GACTCCTGAC	960
40	AGCCTCCTGC ACCTGTGCAA GGGAACTGTG GGGACGCACG AGGATGCCCC CCARGGCCTT	1020
	GGGGAAAAGC CCCCACTGCC CCTCACTCTT CTCTGGACCC CCACCCTCCA TCCTCACCCA	1080
45	GCTCCCGGGG GTGGGGTCGG GTGAGGGCAG CAGGGATGCC CGCCAGGGAC TTGCAAGGAC	1140
75	CCCCTGGGTT TTGAGGGTGT CCCATTCTCA ACTCTAATCC ATCCCAGCCC TCTGGAGGAT	1200
	TTGGGGTGCC CCTCTCGGCA GGGAACAGGA AGTAGGAATC CCAGAAGGGT CTGGGGGAAC	1260
50	CCTAACCCTG AGCTCAGTCC AGTTCACCCC TCACCTCCAG CCTGGGGGTC TCCAGACACT	1320
	GCCAGGGCCC CCTCAGGACG GCTGGAGCCT GGAGGAGACA GCCACGGGGT GGTGGGCTGG	1380
55	GCCTGGACCC CACCGTGGTG GGCAGCAGGG CTGCCCGGCA GGCTTGGTGG ACTCTGCTGG	1440
23	CAGCAAATAA AGAGATGACG GCAAAAAAAA AAAAAAAAAA	1500
	AAAAAAAAA AAACCCACCG TCCGC	1525

(2) INFORMATION FOR SEQ ID NO: 80:

5	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1563 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	·
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 80:	
	AATTCGGCAC GAGNCAGAAA CCTGCGGAAA ATGGTAGCGA TGGCGGCTGG GCCGAGTGGG	60
15	TGTCTGGTGC CGGCGTTTGG GCTACGGTTG TTGTTGGCGA CTGTGCTTCA AGCGGTGTCT	120
	GCTTTTGGGG CAGAGTTTTC ATCGGAGGCA TGCAGAGAGT TAGGCTTTTC TAGCAACTTG	180
20	CTTTGCAGCT CTTGTGATCT TCTCGGACAG TTCAACCTGC TTCAGCTGGA TCCTGATTGC	240
20	AGAGGATGCT GTCAGGAGGA AGCACAATTT GAAACCAAAA AGCTGTATGC AGGAGCTATT	300
	CTTGAAGTTT GTGGATGAAA ATTGGGAAGG TTCCCTCAAG TCCAAGCTTT TGTTAGGAGT	360
25	GATAAACCCA AACTGTTCAG AGGACTGCAA ATCAAGTATG TCCGTGGTTC AGACCCTGTA	420
	TTAAAGCTTT TGGACGACAA TGGGAACATT GCTGAAGAAC TGAGCATTCT CAAATGGAAC	480
30	ACAGACAGTG TAGAAGAATT CCTGAGTGAA AAGTTGGAAC GCATATAAAT CTTGCTTAAA	540
20	TTTTGTCCTA TCCTTTTGTT ACCTTATCAA ATGAAATATT ACAGCACCTA GAAAATAATT	600
	TAGTTTTGCT TGCTTCCATT GATCAGTCTT TTACTTGAGG CATTAAATAT CTAATTAAAT	660
35	CGTGAAATGG CAGTATAGTC CATGATATCT AAGGAGTTGG CAAGCTTAAC AAAACCCATT	720
	TTTTATAAAT GTCCATCCTC CTGCATTTGT TGATACCACT AACAAAATGC TTTGTAACAG	780
40	ACTTGCGGTT AATTATGCAA ATGATAGTTT GTGATAATTG GTCCAGTTTT ACGAACAACA	840
	GATTTCTAAA TTAGAGAGGT TAACAAGACA GATGATTACT ATGCCTCATG TGCTGTGTGC	900
	TCTTTGAAAG GAATGACAGC AGACTACAAA GCAAATAAGA TATACTGAGC CTCAACAGAT	960
45	TGCCTGCTCC TCAGAGTCTC TCCTATTTTT GTATTACCCA GCTTTCTTTT TAATACAAAT	1020
	GTTATTTATA GTTTACAATG AATGCACTGC ATAAAAACTT TGTAGCTTCA TTATTGTAAA	1080
50	ACATATTCAA GATCCTACAG TAAGAGTGAA ACATTCACAA AGATTTGCGT TAATGAAGAC	1140
	TACACAGAAA ACCTTTCTAG GGATTTGTGT GGATCAGATA CATACTTGGC AAATTTTTGA	1200
٠	GTTTTACATT CTTACAGAAA AGTCCATTTA AAAGTGATCA TTTGTAAGAC CAAAATATAA	1260
55	ATAAAAAGTT TCAAAAATCT ATCTGAATTT GGAATTCTTC TGGTTTGTTC TTTCATGTTT	1320
	AAAAATGATG TITTTCAATG CATTITTTC ATGTAAGCCC TTTTTTTAGC CAAAATGTAA	1380
60	AAATGGCTGT AATATTTAAA ACTTATAACA TCTTATTGTT GGTAATAGTG CTTTATATTT	1440

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	GTCTGATTTT	ATTTTTCAAA	GTTTTTTCAT	TTATGAACAC	ATTTTCATTG	GTATATTATT	1500
	TAAGGAATAT	CTCTTGATAT	AGAATTTTTA	TATTAAAAAT	GATTTTTCTT	TGCTTAAAAA	1560
5	AAA						1563

10 (2) INFORMATION FOR SEQ ID NO: 81:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1020 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 81:

20	TGCACGCTGG	CCATGTGGGN	GTTGGGCCAC	TGCGACCCCC	GGCGCTGCAC	GGGCCGCAAG	60
	CTGGCCCGCC	TGGGGCTGGT	GCGCTGCCTG	CGCCTGGGCC	ACAGATTCGG	CGGTCTGGTG	120
25	CTGAGCCCCG	TGGGCAAGCA	GTACGCGTCC	CCCGCAGACA	GACAGCTGGT	GGCGCAGTCT	180
25	GGGGTCGCCG	TCATCGACTG	CTCCTGGGCC	AGGCTGGACG	AGACACCGTT	TGGGAAGATG	240
	CGAGGGAGCC	ACTTGCGCCT	GTTGCCCTAC	CTGGTGGCCG	CCAACCCCGT	GAACTATGGC	300
30	CGGCCCTACA	GACTTTCCTG	CGTGGAAGCG	TTTGCTGCCA	CCTTCTGCAT	CGTAGGCTTT	360
	CCAGACCTTG	CTGTCATTTT	GCTGCGGAAG	TTTAAATGGG	GCAAGGGCTT	CTTGGACCTG	420
25	AACCGCCAGC	TCCTGGACAA	GTACGCGGCC	TGCGGCAGCC	CGGAGGAGGT	GCTGCAGGCG	480
35	GAGCAGGAGT	TCTTGGCCAA	TGCCAAGGAG	AGCCCCCAGG	AGGAGGAGAT	CGATCCCTTC	540
	GATGTGGATT	CAGGGAGAGA	GTTTGGAAAC	CCCAACAGGC	CTGTGGCCAG	CACCCGGCTG	600
40	CCCTCGGACA	CTGATGACAG	TGATGCGTCT	GAGGACCCAG	GGCCTKGCGC	CGAGCGCGGA	660
	GGAGCCAGCA	GCAGCTGCTG	TGAAGAGGAG	CAGACGCAGG	GACGGGGGC	TGAGGCCAGG	720
4.5	GCCCCGGCTG	AGGTTTGGAA	AGGAATCAAG	AAACGGCAGA	GAGACTGAGG	GTTGCAGACA	780
45	CATATATTTT	TGAGGCTGGG	TGACGAGAAA	ATCTAGAGAC	ATGAGGGACA	TAAATGGGCC	840
	TGGCAGCCTC	GGCTCTTTGC	GGCTGCTGGC	AGGACTGAGC	TGTCCGGGTT	CTCCCCACAC	900
50	TTCCAGCACA	GCTGTGCTCT	GTGTCCTGCC	TCGGCGCTCT	' CGCAAATGAA	GCTGCAGGCC	960
	AAGAAAAAA	AAAAAAAA	AAAAAAAA	AAAAAAAA	AAAAAAAAA	GGGGGGGGC	1020

55

(2) INFORMATION FOR SEQ ID NO: 82:

(i) SEQUENCE CHARACTERISTICS:

60 (A) LENGTH: 770 base pairs

(B) TYPE: nucleic acid

	(C) STRANDEDNESS: double (D) TOPOLOGY: linear					
5	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 82:					
	TCGACCCACG CGTCCGGGCC GCCGTAGCGC GTCTTGGGTC TCCCGGCTGC CGCTGCTGCC	60				
10	GCCGCCGCCT CGGGTCGTGG AGCCAGGAGC GACGTCACCG CCATGGCAGG CATCAAAGCT	120				
10	TTGATTAGTT TGTCCTTTGG AGGAGCAATC GGACTGATGT TTTTRATGCT TGGATGTGCC	180				
	CTTCCAATAT ACAACAAATA CTGGCCCCTC TTTGTTCTAT TTTTTTACAT CCTTTCACCT	240				
15	ATTCCATACT GCATAGCAAG AAGATTAGTG GATGA NACAG ATGCTATGAG TAACGCTTGT	300				
	AAGGAACTTG CCATCTTTCT TACAACGGGC ATTGTCGTGT CAGCTTTTGG ACTCCCTATT	360				
20	GTATTTGCCA GAGCACATCT GATTGAGTGG GGAGCTTGTG CACTTGTTCT CACAGGAAAC	420				
20	ACAGTCATCT TTGCAACTAT ACTAGGCTTT TTCTTGGTCT TTGGAAGCAA TGACGACTTC	480				
	AGCTGGCAGC AGTGGTGAAA AGAAATTACT GAACTATTGT CAAATGGACT TCCTGTCATT	540				
25	TGTTGGCCAT TCACGCACAC AGGAGATGGG GCAGTTAATG CTGAATGGTA TAGCAAGCCT	600				
	CTTGGGGGTA TTTTAGGTGC TCCCTTCTCA CTTTTATTGT AAGCATACTA TTTTCACAGA	660				
30	GACTTGCTGA AGGATTAAAA GGATTTTCTC TTTTGGAAAA AAAAAAAAAA	720				
50	GGGGGGCCC GTWCCCATTC SCCCYATATG AATTCCNTTT TTACAATCCC	770				
35	(2) INFORMATION FOR SEQ ID NO: 83:					
40	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 481 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear					
45	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 83:					
	GAATTCGGCA CGAGCATAGT GTTAACCACT AGAATTCACT GCCCTTCCTA TCCAAAAATG	60				
	ACACTACTGA TCATTTTTCT TCCTTTTSCT TTTACAACAT TMACAAATTC AGGTGGCTCT	120				
50	TTCCCAGTAC GGTAGGCTGA TTCGTATGGA TGCACCACGG TTGGTGACTC CCCCCACCCC	180				
	ACAGAGTTTC TGGCGTTCAT TCGGTTGAAC CCAAGGCCAG CAAGGGCTGA CTGGGAACAA	240				
55	ACCGAACACT AGGCCGTGAA CCAATCGTCT CTCCGTGCCC GGGAGCGAMC CCGGGGGCCT	300				
55	TTCACTCTCC CAAGGACTCC ANGGGGGGGC CGGGTACCCA ATTCCGCCCC TATAGTGAAT	360				
	CCGTNATTAC AATTCCACNT GGGCCGTCCN TTTTTACAAA CGTTCCGTTG AACTGGGAAA	420				
60	AACCCCTTGG CGGTTTACCC CAACTTTAAT CCGCCTTTGC AAGCACATCC CCCCCCTTTT	480				

	С	481
5		
J	(2) INFORMATION FOR SEQ ID NO: 84:	
10	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 644 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 84:	
	GCTGGGATAG AGCATGAAAG GAGAACTGCT CCCTTTTCTG TTTCTCACAG TTTGGTTATG	60
20	GCTTTATAAA CTTKTATTTG GTGAAAGCCC CAGATACCCA AATGTCATTG GCAAAACTTA	120
20	TTTTTTTTTC TGGACAGATC AGATTTCTAG AGAGAGCAGA TTTCTAGAGA GATTAGCATT	180
	CATAGTAAGT GAAAATTGTC TAATTTTTT AATCCATGCT ATTACTGGGC AGTAGGTCTA	240
25	ATTTTTTTG ACAAAAATA GATCTATTTT CCTTATATAT TGATTTAGAA TCTTAAGTTA	300
	GAATTTTATA GAAGAAATGT CTGAGCAGTT CTATGTATGG AGGAGCAATT CAGCTTTTCA	360
30	GCAGCAACTT TATCTTTTGC CACTAGAGGG AGATCTGTGG TTGCTTTCTC CTTTGGAGAA	420
30	TAGCTGCTTT GCTTTTATTT TTAATTTCTA AGGTTGGAAT AGAACTTATT CTCAAAATTC	480
	CTTTAGTGTT ATTAAATATT TTCATTTATT AGTCAAAGGT AAGTTAATTA AGCTTGTTTA	540
35	ATGATGCCAA TCTTATGCTT TTCTGTAATC TTCAATTTTT AATAAATGTG AGTTAGATAC	600
	TAAGTGAAAA AAAAAAAAAAAAAAAAAAAAAAAAAAAA	644
40		
	(2) INFORMATION FOR SEQ ID NO: 85:	
45	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1351 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 85:	
	GGCACGAGTG CGCASGCGTG GGGCTCTCTC CTTGTCAGTC GGCGCCGCGT GCGGGCTGGT	60
55	GGCTCTGTGG CAGCGGCGGC GGCAGGACTC CGGCACTATG AGCGGCTTCA GCACCGAGGA	120
33	GCGCGCCGCG CCNTTCTCCC TGGAGTACCG AGTCTTCCTC AAAAATGAGA AAGGACAATA	180
	TATATCTCCA TTTCATGATA TTCCAATTTA TGCAGATAAG GATGTGTTTC ACATGGTAGT	240
60	THE ACTION CONTROL AND AND AND CONTRACT ACADAGGAC CONTRACT	300

	TATTAAACAA	GATGTGAAAA	AAGGAAAACT	TCGCTATGTT	GCGAATTTGT	TCCCGTATAA	360
5	AGGATATATC	TGGAACTATG	GTGCCATCCC	TCAGACTTGG	GAAGACCCAG	GGCACAATGA	420
J	TAAACATACT	GGCTGTTGTG	GTGACAATGA	CCCAATTGAT	GTGTGTGAAA	TTGGAAGCAA	480
	GGTATGTGCA	AGAGGTGAAA	TAATTGGCGT	GAAAGTTCTA	GGCATATTGG	CTATGATTGA	540
10	CGAAGGGGAA	ACCGACTGGA	AAGTCATTGC	CATTAATGTG	GATGATCCTG	ATGCAGCCAA	600
	TTATAATGAT	ATCAATGATG	TCAAACGGCT	GAAACCTGGC	TACTTAGAAG	CTACTGTGGA	660
15	CTGGTTTAGA	AGGTATAAGG	TTCCTGATGG	AAAACCAGAA	AATGAGTTTG	CGTTTAATGC	720
13	AGAATTTAAA	GATAAGGACT	TTGCCATTGA	ТАТТАТТААА	AGCACTCATG	ACCATTGGAA	780
	AGCATTAGTG	ACTAAGAAAA	CGAATGGAAA	AGGAATCAGT	TGCATGAATA	CAACTTTGTC	840
20	TGAGAGCCCC	TTCAAGTGTG	ATCCTGATGC	TGCCAGAGCC	ATTGTGGATG	CTTTACCACC	900
	ACCCTGTGAA	TCTGCCTGCA	CAGTACCAAC	AGACGTGGAT	AAGTGGTTCC	ATCACCAGAA	960
25	AAACTAATGA	GATTTCTCTG	GAATACAAGC	TGATATTGCT	ACATCGTGTT	CATCTGGATG	1020
23	TATTAGAAGT	AAAAGTAGTA	GCTTTTCAAA	GCTTTAAATT	TGTAGAACTC	АТСТААСТАА	1080
	AGTAAATTCT	GCTGTGACTA	ATCCAATATA	CTCAGAATGT	TATCCATCTA	AAGCATTTTT	1140
30	CATATCTCAA	CTAAGATAAC	TTTTAGCACA	TGCTTAAATA	TCAAAGCAGT	TGTCATTTGG	1200
	AAGTCACTTG	TGAATAGATG	TGCAAGGGGA	GCACATATTG	GATGTATATG	TTACCATATG	1260
35	TTAGGAAATA	AAATTATTTT	GCTGAAAAA	АААААААА	AACCNCGGGG	GGGGCCCCGG	1320
55	TCCCCATTTG	GCCCTTTGGG	GGGNGGTTTT	A			1351
40	(2) INFORMA	ATION FOR SI	EQ ID NO: 86	5:			
45	(i)	(B) TYP (C) STR	HARACTERIST GTH: 2527 b E: nucleic ANDEDNESS: OLOGY: line	ase pairs acid double			
50	(xi) SEQUENCE	DESCRIPTION	: SEQ ID NO	: 86:		
50	CTCTTGCTAC	CTTCCCGGCG	CAGAGAACCC	CGGCTGCTCA	GCGCGCTCCG	GGGTCATGGA	60
	GATCCCCGGG	AGCCTGTGCA	AGAAAGTCAA	GCTGAGCAAT	AACGCGCAGA	ACTGGGGAAT	120
55	GCAGAGAGCA	ACCAATGTCA	CCTACCAAGC	CCATCATGTC	AGCAGGAACA	AGAGAGGTCA	180
	GGTGGTGGGG	ACCAGAGGTG	GCTTTCGTGG	TTGCACAGTT	TGGCTAACAG	GCTTGTCTGG	240
60	AGCGGGAAAG	ACTACTGTGA	GCATGGCCTT	GGAGGAGTAC	CTGGTTTGTC	ATGGTATTCC	300

	ATGCTACACT	CTGGATGGTG	ACAATATTCG	TCAAGGTCTC	AATAAAAATC	TTGGCTTTAG	360
	TCCTGAAGAC	AGAGAAGAGA	ATGTTCGACG	CATCGCAGAA	GTTGCTAAAC	TGTTTGCAGA	420
5	TGCTGGCTTA	GTGTGCATCA	CAAGTTTCAT	ATCACCTTAC	ACTCAGGATC	GCAACAATGC	480
	AAGGCAAATT	CATGAAGGTG	CAAGTTTACC	GTTTTTTGAA	GTATTTGTTG	ATGCTCCTCT	540
10	GCATGTTTGT	GAACAGAGGG	ATGTCAAAGG	ACTCTACAAA	AAAGCCCGGG	CAGGAGAAAT	600
10	TAAAGGTTTC	ACTGGGATCG	ATTCTGAATA	TGAAAAGCCA	GAGGCCCCTG	AGTTGGTGCT	660
	GAAAACAGAC	TCCTGTGATG	TAAATGACTG	TGTCCAGCAA	GTTGTGGAAC	TTCTACAGGA	720
15	ACGGGATATT	GTACCTGTGG	ATGCATCTTA	TGAAGTAAAA	GAACTATATG	TGCCAGAAAA	780
	TAAACTTCAT	TTGGCAAAAA	CAGATGCGGA	AACATTACCA	GCACTGAAAA	TTAATAAAGT	840
20	GGATATGCAG	TGGGTGCAGG	TTTTGGCAGA	AGGTTGGGCA	ACCCCATTGA	ATGGCTTTAT	900
20	GAGAGAGAGG	GAGTACTTGC	AGTGCCTTCA	TTTTGATTGT	CTTCTGGATG	GAGGTGTCAT	960
	TAACTTGTCA	GTACCTATAG	TTCTGACTGC	GACTCATGAA	GATAAAGAGA	GGCTGGACGG	1020
25	CTGTACAGCA	TTTGCTCTGA	TGTATGAGGG	CCGCCGTGTG	GCCATTCTTC	GCAATCCAGA	1080
	GTTTTTTGAG	CACAGGAAAG	AGGAGCGCTG	TGCCAGACAG	TGGGGAACGA	CATGCAAGAA	1140
30	CCACCCCTAT	ATTAAGATGG	TGATGGAACA	AGGAGATTGG	CTGATTGGAG	GAGATCTTCA	1200
50	AGTCTTGGAT	CGAGTTTATT	GGAATGATGG	TCTTGATCAG	TATCGTCTTA	CTCCTACTGA	1260
	GCTAAAGCAG	AAATTTAAAG	ATATGAATGC	TGATGCTGTC	TTTGCATTTC	AACTACGCAA	1320
35	CCCAGTGCAC	AATGGACATG	CCCTGTTAAT	GCAGGATACC	CATAAGCAAC	TTCTAGAGAG	1380
	GGGCTACCGG	CGCCCTGTCC	TCCTCCTCCA	CCCTCTGGGT	GGCTGGACAA	AGGATGACGA	1440
40	TGTTCCTTTG	ATGTGGCGTA	TGAAGCAGCA	TGCTGCAGTG	TTGGAGGAAG	GAGTTCTGAA	1500
	TCCTGAGACG	ACAGTGGTGG	CCATCTTCCC	ATCTCCCATG	ATGTATGCTG	GACCAACTGA	1560
	GGTCCAGTGG	CATTGCAGAG	CACGGATGGT	TGCAGGAGCC	AACTTTTACA	TTGTTGGACG	1620
45	AGACCCTGCT	GCATGCCTC	ATCCAGAAAC	AGGGAAGGAT	CTTTATGAGC	CAAGTCATGG	1680
	TGCCAAAGTG	CTGACGATGG	CCCCTGGTTT	AATCACTTTG	GAAATAGTTC	CCTTTCGAGT	1740
50	TGCAGCTTAC	: AACAAGAAAA	AGAAGCGTAT	GGACTACTAI	GACTCTGAAC	ACCATGAAGA	1800
	CTTTGAATTT	· ATTTCAGGAA	CACGAATGCG	CAAACTTGCT	CGAGAAGGCC	AGAAACCACC	1860
	TGAAGGTTTC	ATGGCTCCC#	AGGCTTGGAC	CGTGCTGACA	A GAATACTACA	AATCCTTGGA	1920
55	GAAAGCTTAG	GCTGTTAACC	CAGTCACTCC	: ACCTTTGACA	A CATTACTAGI	' AACAAGAGGG	1980
	GACCACATAC	TCTCTGTTGC	CATTTCTTTC	TGGTGTCTG	r ctggacatgo	TTCCTAAAAA	2040
60	CAGACCATT	TCCTTAACT	GCATCAGTTI	TGGTCTGCCT	TATGAGTTCT	GTTTTGAACA	2100

	AGTGTAACAC ACTGATGGTT TTAATGTATC TTTTCCACTT ATTATAGTTA TATTCCTACA	2160
	ATACAATTTT AAAATTGTCT TTTTATATTA TATTTATGCT TCTGTGTCAT GATTTTTTCA	2220
5	AGCTGTTATA TTAGTTGTAA CCAGTAGTAT TCACATTAAA TCTTGCTTTT TTTCCCCTTA	2280
	AAAAAAGAAA AAAATTACCA AACAATAAAC TTGGCTAGAC CTTGTTTTGA GGATTTTACA	2340
10	AGACCTTTGT AGCGATTAGA TTTTTTTTCT ACATTGAAAA TAGAAACTGC TTCCTTTCTT	2400
10	CTTTCCAGTC AGCTATTGGT CTTTCCAGCT GTTATAATCT AAAGTATTCT TATGATCTGT	2460
	GTAAGCTCTG AATGAACTTC TTTACTCAAT AAAATTAATT TTTTGGCTTC TTAAAAAAAA	2520
15	алалала	2527
20	(2) INFORMATION FOR SEQ ID NO: 87:	
25	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 2566 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 87:	
30	CCCAAGAATT CGGCACGAGC GNGGCAWAAK TGGGATTTCT GAAACCTGTA GGCCCCAAGC	60
	CCATCAACTT GCCCAAAGAA GATTCCAAAC CTACATTTCC CTGGCCTSCT GGAAACAAGC	120
35	CATCTCTTCA CAGTGTAAAC CAAGACCATG ACTTAAAGCC ACTAGGCCGA AATCTGGGCC	180
55	TACTCCTCCA ACCTCAGAAA ATGAACAGAA GCAAGCKTTT CCCAAATTGA CTGGGGTTAA	240
	AGGGAAATTT ATGTCAGCAT CACAAGATCT TGAACCCAAG CCCCTCTTCC CCAAACCCGC	300
40	CTTTGGCCAG AAGCCGCCCC TAAGTACCGA GAACTCCCAT GAAGACGAAA GCCCCATGAA	360
	GAATGTGTCT TCATCAAAAG GGTCCCCAGC TCCCCTGGGA GTCAGGTCCA AAAGCGGCCC	420
45	TTTAAAACCA GCAAGGGAAG ACTCAGAAAA TAAAGACCAT GCAGGGGAGA TTTCAAGTTT	480
	GCCCTTTCCT GGAGTGGTTT TGAAACCTGC TGCGAGCAGG GGAGGCCCAG GTCTCTCCAA	540
	AAATGGTGAA GAAAAAAAGG AAGATAGGAA GATAGATGCT GCTAAGAACA CCTTCCAGAG	600
50	CAAAATAAAT CAGGAAGAGT TGGCCTCAGG GACTCCTCCT GCCAGGTTCC CTAAGGCCCC	660
	TTCTAAGCTG ACAGTGGGGG GGCCATGGGG CCAAAGTCAG GAAAAGGAAA AGGGAGACAA	720
55	GAATTCAGCC ACCCCGAAAC AGAAGCCATT GCCTCCCTTG TTTACCTTGG GTCCACCTCC	780
•	ACCAAAACCC AACAGACCAC CAAATGTTGA CCTGACGAAA TTCCACAAAA CCTCTTCTGG	840
	AAACAGTACT AGCAAAGGCC AGACGTCTTA CTCAACAACT TCCCTGCCAC CACCTCCACC	900
60	ATCCCATCCG GCCAGCCAAC CACCATTGCC AGCATCTCAC CCATCACAAC CACCAGTCCC	960

	AAGCCTACCT	CCCAGAAACA	TTAAACCTCC	GTTTGACCTA	AAAAGCCCTG	TCAATGAAGA	1020
_	CAATCAAGAT	GGTGTCACGC	ACTCTGATGG	TGCTGGAAAT	CTAGATGAGG	AACAAGACAG	1080
5	TGAAGGAGAA	ACATATGAAG	ACATAGAAGC	ATCCAAAGAA	AGAGAGAAGA	AAAGGGAAAA	1140
	GGAAGAAAAG	AAGAGGTTAG	AGCTGGAGAA	AAAGGAACAG	AAAGAGAAAG	AAAAGAAAGA	1200
10	ACAAGAAATA	AAGAAGAAAT	TTAAACTAAC	AGGCCCTATT	CAAGTCATCC	ATCTTGCAAA	1260
	AGCTTGTTGT	GATGTCAAAG	GAGGAAAGAA	TGAACTGAGC	TTCAAGCAAG	GAGAGCAAAT	1320
15	TGAAATCATC	CGCATCACAG	ACAACCCAGA	AGGAAAATGG	TTGGGCAGAA	CAGCAAGGGG	1380
15	TTCATATGGC	TATATTAAAA	CAACTGCTGT	AGAGATTGAC	TATGATTCTT	TGAAACTGAA	1440
	AAAAGACTCT	CTTGGTGCCC	CTTCAAGACC	TATTGAAGAT	GACCAAGAAG	TATATGATGA	1500
20	TGTTGCAGAG	CAGGATGATA	TTAGCAGCCA	CAGTCAGAGT	GGAAGTGGAG	GGATATTCCC	1560
	TCCACCACCA	GATGATGACA	TTTATGATGG	GATTGAAGAG	GAAGATGCTG	ATGATGGCTC	1620
25	CACACTACAG	GTTCAAGAGA	AGAGTAATAC	GTGGTCCTGG	GGGATTTTGA	AGATGTTAAA	1680
23	GGGAAAAGAT	GACAGAAAGA	AAAGTATACG	AGAGAAACCT	AAAGTCTCTG	ACTCAGACAA	1740
	TAATGAAGGT	TCATCTTTCC	CTGCTCCTCC	TAAACAATTG	GACATGGGAG	ATGAAGTTTA	1800
30	CGATGATGTG	GATACCTCTG	ATTTCCCTGT	TTCATCAGCA	GAGATGAGTC	AAGGAACTAA	1860
	TGTTGGAAAA	GCTAAGACAG	AAGAAAAGGA	CCTTAAGAAG	CTAAAAAAGC	AGRAAAAARA	1920
35	ARAAAAAGAC	TTCAGGAAAA	ATTTAAATA	TGATGGTGAA	ATTAGAGTCC	TATATTCAAC	1980
55	TAAAGTTACA	ACTTCCATAA	CTTCTAAAAA	GTGGGGAACC	AGAGATCTAC	AGGTAAAACC	2040
	TGGTGAATCT	CTAGAAGTTA	TACAAACCAC	AGATGACACA	A AAAGTTCTCT	GCAGAAATGA	2100
40	AGAAGGGAAA	TATGGTTATG	TCCTTCGGAG	TTACCTAGCO	GACAATGATG	GAGAGATCTA	2160
	TGATGATATT	GCTGATGGCT	GCATCTATGA	CAATGACTAC	CACTCAACTT	TGGTCATTCT	2220
45	GCTGTGTTCA	TTAGGTGCCA	ATGTGAAGTC	TGGATTTA	A TTGGCATGTT	ATTGGGTATC	2280
10	AAGAAAATTA	ATGCACAAA	A CCACTTATTA	A TCATTTGTT	A TGAAATCCCA	A ATTATCTTTA	2340
	CAAAGTGTTT	AAAGTTTGAA	A CATAGAAAAT	AATCTCTCT(G CTTAATTGTT	ATCTCAGAAG	2400
50	ACTACATTAC	TGAGATGTA	A GAATTATTA	A ATATTCCAT	r TCCGCTTTGC	G CTACAATTAT	2460
	GAAGAAGTTO	AAGGTACTT	C TTTTAGACCA	A CCAGTAAAT.	A ATCCTCCTTC	AAAAAATAAA	2520
55	AAAAAATAA	AAAAAAAA	A ACTCGAGGG	GGCCCGGT.	A CCCAAT		2566

⁽²⁾ INFORMATION FOR SEQ ID NO: 88:

5	(A) LENGTH: 540 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 88:	
10	GAATTCGGCA CGAGGCTTTC TGTGTCCTCT GTGGCTGCTT TAGTGTGCCA CCAGGGGCAG	60
10	ACTIGGGTGG GTTGCAGCAG AGATGGCATG GCCCTCAAGG TCCAAGATGT TTACTCTCTT	120
	GCCGGTCCTC TGTTATCTCT GGTCTTTGTG GTTGCCACAG TTTTCTTGGA TCCAGGAGTT	180
15	AAAGGCAGTC CTGAGGGATG ATGGCCTCAT CTCCGCAGTT GCYTGGAATG CTGAATTTCA	240
	GACGTGCTAA AGGAGGGTTG CAGACATTGT GTGGWATGCA TTCAGACCCC AGATGTGGGT	300
20	GCAGGAAGGC AGGCATGGCA CAGCCAGGTA GAGACTGGTT TCCAGGCCCA AGCAGCCTTC	360
_ •	AGCAGCTGTG CGCCTTGTTT CTGATGTTGT TTGGGAGTAA GAATAATGTA GACATGGGGG	420
	GTCATGARGC TCAATAAAAA CTTCAAGGAA ACCTCCCATG GCATGGTTGG GCGCAGTGAC	480
25	TCATGCCTGT AACCCCAGCA CTGTGGAATG CCAAGGTGGA AGGATCGCTT GAGGCCAAGA	540
30	(2) INFORMATION FOR SEQ ID NO: 89:	•
35	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1863 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
35	(A) LENGTH: 1863 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double	
35 40	(A) LENGTH: 1863 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	60
	 (A) LENGTH: 1863 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 89: 	60 120
40	(A) LENGTH: 1863 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 89: TCGACCCACG CGTCCGGCGA GATCCCTACC GCAGTAGCCG CCTCTGCCGC CGCGGAGCTT	
	(A) LENGTH: 1863 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 89: TCGACCCACG CGTCCGGCGA GATCCCTACC GCAGTAGCCG CCTCTGCCGC CGCGGAGCTT CCCGAACCTC TTCAGCCGCC CGGAGCCGCT CCCGGAGCCC GGCCGTAGAG GCTGCAATCG	120
40	(A) LENGTH: 1863 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 89: TCGACCCACG CGTCCGGCGA GATCCCTACC GCAGTAGCCG CCTCTGCCGC CGCGGAGCTT CCCGAACCTC TTCAGCCGCC CGGAGCCGCT CCCGGAGCCC GGCCGTAGAG GCTGCAATCG CAGCCGGGAG CCCGCAGCCC GCGCCCCGAG CCCGCCGCGC CCCTTCGAGG GCGCCCCAGG	120 180
40	(A) LENGTH: 1863 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 89: TCGACCCACG CGTCCGGCGA GATCCCTACC GCAGTAGCCG CCTCTGCCGC CGCGGAGCTT CCCGAACCTC TTCAGCCGCC CGGAGCCGCT CCCGGAGCCC GGCCGTAGAG GCTGCAATCG CAGCCGGGAG CCCGCAGCCC GCGCCCCGAG CCCGCCGCGC CCCTTCGAGG GCGCCCCAGG CCGCGCCCATG GTGAAGGTGA CGTTCAACTC CGCTCTGGCC CAGAAGGAGG CCAAGAAGGA	120 180 240
40 45	(A) LENGTH: 1863 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 89: TCGACCCACG CGTCCGGCGA GATCCCTACC GCAGTAGCCG CCTCTGCCGC CGCGGAGCTT CCCGAACCTC TTCAGCCGCC CGGAGCCGCT CCCGGAGCCC GGCCGTAGAG GCTGCAATCG CAGCCGGGAG CCCGCAGCCC GCGCCCCGAG CCCGCCGCCG CCCTTCGAGG GCGCCCCAGG CCGCGCCCATG GTGAAGGTGA CGTTCAACTC CGCTCTGGCC CAGAAGGAGG CGAGCCCAAG AGCGGCGAGG AGGCGCTCAT CATCCCCCCC GACGCCGTCG CGGTGGACTG	120 180 240 300
40 45 50	(A) LENGTH: 1863 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 89: TCGACCCACG CGTCCGGCGA GATCCCTACC GCAGTAGCCG CCTCTGCCGC CGCGGAGCTT CCCGAACCTC TTCAGCCGCC CGGAGCCGCT CCCGGAGCCC GGCCGTAGAG GCTGCAATCG CAGCCGGGAG CCCGCAGCCC GCGCCCCGAG CCCGCCGCCG CCCTTCGAGG GCGCCCCAGG CCGCGCCATG GTGAAGGTGA CGTTCAACTC CGCTCTGGCC CAGAAGGAGG CCAAGAAGGA CGAGCCCAAG AGCGGCGAGG AGGCGCTCAT CATCCCCCCC GACGCCGTCG CGGTGGACTG CAAGGACCCA GATGATGTGG TACCAGTTGG CCAAAGAAGA GCCTGGTGTT GGTGCATGTG	120 180 240 300 360
40 45	(A) LENGTH: 1863 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 89: TCGACCCACG CGTCCGGCGA GATCCCTACC GCAGTAGCCG CCTCTGCCGC CGCGGAGCTT CCCGAACCTC TTCAGCCGCC CGGAGCCGCT CCCGGAGCCC GGCCGTAGAG GCTGCAATCG CAGCCGGGAG CCCGCAGCCC GCGCCCCGAG CCCGCCGCCG CCCTTCGAGG GCGCCCCAGG CCGCGCCATG GTGAAGGTGA CGTTCAACTC CGCTCTGGCC CAGAAGGAGG CCAAGAAGGA CGAGCCCAAG AGCGGCGAGG AGGCGCTCAT CATCCCCCCC GACGCCGTCG CGGTGGACTG CAAGGACCCA GATGATGTGG TACCAGTTGG CCAAAGAAGA GCCTGGTGTT GGTGCATGTG CTTTGGACTA GCATTTATGC TTGCAGGTGT TATTCTAGGA GGAGCATACT TGTACAAATA	120 180 240 300 360 420
40 45 50	(A) LENGTH: 1863 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 89: TCGACCCACG CGTCCGGCGA GATCCCTACC GCAGTAGCCG CCTCTGCCGC CGCGGAGCTT CCCGAACCTC TTCAGCCGCC CGGAGCCGCT CCCGGAGCCC GGCCGTAGAG GCTGCAATCG CAGCCGGGAG CCCGCAGCCC GCGCCCCGAG CCCGCCGCCG CCCTTCGAGG GCGCCCCAGG CCGCGCCATG GTGAAGGTGA CGTTCAACTC CGCTCTGGCC CAGAAGGAGG CCAAGAAGGA CGAGCCCAAG AGCGGCGAGG AGGCGCTCAT CATCCCCCCC GACGCCGTCG CGGTGGACTG CAAGGACCCA GATGATGTGG TACCAGTTGG CCAAAGAAGA GCCTGGTGTT GGTGCATGTG CTTTGGACTA GCATTTATGC TTGCAGGTGT TATTCTAGGA GGAGCATACT TGTACAAATA TTTTGCACTT CAACCAGATG ACGTGTACTA CTGTGGAATA AAGTACATCA AAGATGATGT	120 180 240 300 360 420 480

	TCTTAACCTG GATAAGTGCT ATGTGATCCC TCTGAACACT TCCATTGTTA TGCCACCCAG	720
	AAACCTACTG GAGTTACTTA TTAACATCAA GGCTGGAACC TATTTGCCTC AGTCCTATCT	780
5	GATTCATGAG CACATGGTTA TTACTGATCG CATTGAAAAC ATTGATCACC TGGGTTTCTT	840
	TATTTATCGA CTGTGTCATG ACAAGGAAAC TTACAAACTG CAACGCAGAG AAACTATTAA	900
10	AGGTATTCAG AAACGTGAAG CCAGCAATTG TTTCGCAATT CGGCATTTTG AAAACAAATT	960
	TGCCGTGGAA ACTTTAATTT GTTCTTGAAC AGTCAAGAAA AACATTATTG AGGAAAATTA	1020
. ~	ATATCACAGC ATAACCCCAC CCTTTACATT TTGTGCAGTG ATTATTTTTT AAAGTCTTCT	1080
15	TTCATGTAAG TAGCAAACAG GGCTTTACTA TCTTTTCATC TCATTAATTC AATTAAAACC	1140
	ATTACCTTAA AATTTTTTC TTTCGAAGTG TGGTGTCTTT TATATTTGAA TTAGTAACTG	1200
20	TATGAAGTCA TAGATAATAG TACATGTCAC CTTAGGTAGT AGGAAGAATT ACAATTTCTT	1260
	TAAATCATTT ATCTGGATTT TTATGTTTTA TTAGCATTTT CAAGAAGACG GATTATCTAG	1320
25	AGAATAATCA TATATATGCA TACGTAAAAA TGGACCACAG TGACTTATTT GTAGTTGTTA	1380
25	GTTGCCCTGC TACCTAGTTT GTTAGTGCAT TTGAGCACAC ATTTTAATTT TCCTCTAATT	1440
	AAAATGTGCA GTATTTTCAG TGTCAAATAT ATTTAACTAT TTAGAGAATG ATTTCCACCT	1500
30	TTATGTTTTA ATATCCTAGG CATCTGCTGT AATAATATTT TAGAAAATGT TTGGAATTTA	1560
	AGAAATAACT TGTGTTACTA ATTTGTATAA CCCATATCTG TGCAATGGAA TATAAATATC	1620
25	ACAAAGTTGT TTAACTAGAC TGCGTGTTGT TTTTCCCGTA TAATAAAACC AAAGAATAGT	1680
35	TTGGTTCTTC AAATCTTAAG AGAATCCACA TAAAAGAAGA AACTATTTTT TAAAAAATTCA	1740
	CTTCTATATA TACAATGAGT AAAATCACAG ATTTTTTCTT TAAATAAAAA TAAGTCATTT	1800
40	TAATAACTAA ACCAGATTCT TTGTGATACT ATTAANGTAA CATTTAGCCC CAAAAAAAA	1860
	AAA	1863
45	(2) INTERPRETARION FOR SEC ID NO. 90.	

(2) INFORMATION FOR SEQ ID NO: 90:

(i) SEQUENCE CHARACTERISTICS: 50

(A) LENGTH: 2478 base pairs (B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 90: 55

> GGCACAGCGG CACGAGGTGA GCTGAGCCGG TGGGTGAGCG GCGGCCACGG CATCCTGTGC 60

TGTGGGGGCT ACGAGGAAAG ATCTAATTAT CATGGACCTG CGACAGTTTC TTATGTGCCT 60

	GTCCCTGTGC	ACAGCCTTTG	CCTTGAGCAA	ACCCACAGAA	AAGAAGGACC	GTGTACATCA	180
	TGAGCCTCAG	CTCAGTGACA	AGGTTCACAA	TGATGCTCAG	AGTTTTGATT	ATGACCATGA	240
5	TGCCTTCTTG	GGTGCTGAAG	AAGCAAAGAC	CTTTGATCAG	CTGACACCAG	AAGAGAGCAA	300
	GGAAAGGCTT	GGAAAGATTG	TAAGTAAAAT	AGATGGCGAC	AAGGACGGGT	TTGTCACTGT	360
10	GGATGAGCTC	AAAGACTGGA	TTAAATTTGC	ACAAAAGCGC	TGGATTTACG	AGGATGTAGA	420
	GCGACAGTGG	AAGGGCATG	ACCTCAATGA	GGACGGCCTC	GTTTCCTGGG	AGGAGTATAA	480
	AAATGCCACC	TACGGCTACG	TTTTAGATGA	TCCAGATCCT	GATGATGGAT	TTAACTATAA	540
15	ACAGATGATG	GTTAGAGATG	AGCGGAGGTT	TAAAATGGCA	GACAAGGATG	GAGACCTCAT	600
	TGCCACCAAG	GAGGAGTTCA	CAGCTTTCCT	GCACCCTGAG	GAGTATGACT	ACATGAAAGA	660
20	TATAGTAGTA	CAGGAAACAA	TGGAAGATAT	AGATAAGAAT	GCTGATGGTT	TCATTGATCT	720
	AGAAGAGTAT	ATTGGTGACA	TGTACAGCCA	TGATGGGAAT	ACTGATGAGC	CAGAATGGGT	780
	AAAGACAGAG	CGAGAGCAGT	TTGTTGAGTT	TCGGGATAAG	AACCGTGATG	GGAAGATGGA	840
25	CAAGGAAGAG	ACCAAAGACT	GGATCCTTCC	CTCAGACTAT	GATCATGCAG	AGGCAGAAGC	900
	CAGGCACCTG	GTCTATGAAT	CAGACCAAAA	CAAGGATGGC	AAGCTTACCA	AGGAGGAGAT	960
30	CGTTGACAAG	TATGACTTAT	TTGTTGGCAG	CCAGGCCACA	GATTTTGGGG	AGGCCTTAGT	1020
	ACGGCATGAT	GAGTTCTGAG	CTRCGGAGGA	ACCCTCATTT	CCTCAAAAGT	AATTTATTTT	1080
	TACAGCTTCT	GGTTTCACAT	GAAATTGTTT	GCGCTACTG:	ACTGTTACT	ACAAACTTTT	1140
35	TAAGACATGA	AAAGGCGTAA	TGAAAACCAT	CCCGTCCCCI	FCCTCCTCC	TCTCTGAGGG	1200
	ACTGGAGGGA	AGCCGTGCTT	CTGAGGAACA	ACTCTAATTA	GTACACTTGT	GTTTGTAGAT	1260
40	TTACACTTTG	TATTATGTAT	TAACATGGCG	TGTTTATTTT	TGTATTTTC	TCTGGTTGGG	1320
	AGTATGATAT	GAAGGATCAA	GATCCTCAAC	TCACACATGT	AGACAAACAT	TAGCTCTTTA	1380
	CTCTTTCTCA	ACCCCTTTTA	TGATTTTAAT	AATTCTCACT	TAACTAATTT	TGTAAGCCTG	1440
45	AGATCAATAA	GAAATGTTCA	GGAGAGAGGA	AAGAAAAAA	ATATATGCTC	CACAATTTAT	1500
	ATTTAGAGAG	AGAACACTTA	GTCTTGCCTG	TCAAAAAGTC	CAACATTTCA	TAGGTAGTAG	1560
50	GGGCCACATA	TTACATTCAG	TTGCTATAGG	TCCAGCAACT	GAACCTGCCA	TTACCTGGGC	1620
	AAGGAAAGAT	CCCTTTGCTC	TAGGAAAGCT	TGGCCCAAAT	TGATTTTCTT	CTTTTTCCCC	1680
	CTGTAGGACT	GACTGTTGGC	TAATTTTGTC	AAGCACAGCT	GTGGTGGGAA	GAGTTAGGGC	1740
55	CAGTGTCTTG	AAAATCAATC	AAGTAGTGAA	TGTGATCTCT	TTGCAGAGCT	ATAGATAGAA	1800
	ACAGCTGGAA	AACTAAAGGA	AAAATACAAG	TGTTTTCGGG	GCATACATTT	TTTTTCTGGG	1860
60	TGTGCATCTG	TTGAAATGCT	CAAGACTTAA	TTATTTGCCT	TTTGAAATCA	CTGTAAATGC	1920

PCT/US98/05311

241

	CCCCATCCGG	TTCCTCTTCT	TCCCAGGTGT	GCCAAGGAAT	TAATCTTGGT	TTCACTACAA	1980
	TTAAAATTCA	CTCCTTTCCA	ATCATGTCAT	TGAAAGTGCC	TTTAACGAAA	GAAATGGTCA	2040
5	CTGAATGGGA	ATTCTCTTAA	GAAACCCTGA	GATTAAAAAA	AGACTATTTG	GATAACTTAT	2100
	AGGAAAGCCT	AGAACCTCCC	AGTAGAGTGG	GGATTTTTTT	CTTCTTCCCT	TTCTCTTTTG	2160
10	GACAATAGTT	AAATTAGCAG	TATTAGTTAT	GAGTTTGGTT	GCAGTGTTCT	TATCTTGTGG	2220
10	GCTGATTTCC	AAAAACCACA	TGCTGCTGAA	TTTACCAGGG	ATCCTCATAC	CTCACAATGC	2280
	AAACCACTTA	CTACCAGGCC	TTTTTCTGTG	TCCACTGGAG	AGCTTGAGCT	CACACTCAAA	2340
15	GATCAGAGGA	CCTACAGAGA	GGGCTCTTTG	GTTTGAGGAC	CATGGCTTAC	CTTTCCTGCC	2400
	TTTGACCCAT	CACACCCCAT	TTCCTCCTCT	TTCCCTCTCC	CCGCTGCCAA	TTCCTGCAGC	2460
20	CCGGGGGAAC	CACTAGTT					2478
20							
	(2) THEODA	INTION FOR S	FO ID NO. 9	1.			
	(2) INFORMATION FOR SEQ ID NO: 91:						

25 (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2058 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

30 (D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 91:

TCGGCCTTGC TTTTGTGGYC TTCCTCTGTG GCCAGAGCGT TTTCATCACC AAGCCTCCTG 60 35 ATGGCAGTNC CTTCACCGAT ATGTTCAAGA TACTGACGTA TTCCTGCTGT TCCCAGAAGC 120 GAAGTGGAGA GCGCCAGAGT AATGGTGAAG GCATTGGAGT NTTTCAGCAA TCTTCTAAAC 180 AAAGTCTGTT TGATTCATGT AAGATGTCTC ATGGTGGGCC ATTTACAGAA GAGAAAGTGG 240 40 AAGATGTGAA AGCTCTGGTC AAGATTGTCC CTGTTTTCTT GGCTTTGATA CCTTACTGGA 300 CAGTGTATTT CCAAATGCAG ACAACATATG TTTTACAGAG TCTTCATTTG AGGATTCCAG 360 45 AAATTTCAAA TATTACAACC ACTCCTCACA CGCTCCCTGC AGCCTGGCTG ACCATGTTTG 420 ATGCTGTGCT CATCCTCCTG CTCATCCCTC TGAAGGACAA ACTGGTCGAT CCCATTTTGA 480 540 GAAGACATGG CCTGCTCCCA TCCTCCCTGA AGAGGATCGC CGTGGGCATG TTCTTTGTCA 50 TGTGCTCRGC CTTTGCTGCA GGAATTTTGG AGAGTAAAAG GCTGAACCTT GTTAAAGAGA 600 AAACCATTAA TCAGACCATC GGCAACGTCG TCTACCATGC TGCCGATCTG TCGCTGTGGT 660 55 GGCAGGTGCC GCAGTACTTG CTGATTGGGA TCAGCGAGAT CTTTGCAAGT ATCGCAGGCC 720 TGGAATTTGC ATACTCAGCT GCCCCCAAGT CCATGCAGAG TGCCATAATG GGCTTGTTCT 780 840 TTTTCTTCTC TGGCGTCGGG TCGTTCGTGG GTTCTGGACT GCTGGCACTG GTGTCTATCA 60

	AAGCCATCGG	ATGGATGAGC	AGTCACACAG	ACTTTGGTAA	TATTAACGGC	TGCTATTTGA	900
5	ACTATTACTT	TITCCTTCTG	GCTGCTATTC	AAGGAGCTAC	CCTCCTGCTT	TTCCTCATTA	960
J	TTTCTGTGAA	ATATGACCAT	CATCGAGACC	ATCAGCGATC	AAGAGCCAAT	GGCGTGCCCA	1020
	CCAGCAGGAG	GGCCTGACCT	TCCTGAGGCC	ATGTGCGGTT	TCTGAGGCTG	ACATGTCAGT	1080
10	AACTGACTGG	GGTGCACTGA	GAACAGGCAA	GACTTTAAAT	TCCCATAAAA	TGTCTGACTT	1140
	CACTGAAACT	TGCATGTTGC	CTGGATTGAT	TTCTTCTTTC	CCTCTATCCA	AAGGAGCTTG	1200
15	GTAAGTGCCT	TACTGCAGCG	TGTCTCCTGG	CACGCTGGGC	CCTCCGGGAG	GAGAGCTGCA	1260
	GATTTCGAGT	ATGTCGCTTG	TCATTCAAGG	TCTCTGTGAA	TCCTCTAGCT	GGGTTCCCTT	1320
	TTTTACAGAA	ACTCACAAAT	GGAGATTGCA	AAGTCTTGGG	GAACTCCACG	TGTTAGTTGG	1380
20	CATCCCAGTT	TCTTAAACAA	ATAGTATCAC	CTGCTTCCCA	TAGCCATATC	TCACTGTAAA	1440
	AAAAAAATT	AATAAACTGT	TACTTATATT	TAAGAAAGTG	AGGATTTTTT	TTTTTTAAAG	1500
25	ATAAAAGCAT	GGTCAGATGC	TGCAAGGATT	TTACATAAAT	GCCATATTTA	TGGTTTCCTT	1560
	CCTGAGAACA	ATCTTGCTCT	TGCCATGTTC	TTTGATTTAG	GCTGGTAGTA	AACACATTTC	1620
	ATCTGCTGCT	TCAAAAAGTA	CTTACTTTTT	AAACCATCAA	CATTACTTTT	CTTTCTTAAG	1680
30	GCAAGGCATG	CATAAGAGTC	ATTTGAGACC	ATGTGTCCCA	TCTCAAGCCA	CAGAGCAACT	1740
	CACGGGGTAC	TTCACACCTT	ACCTAGTCAG	AGTGCTTATA	TATAGCTTTA	TTTTGGTACG	1800
35	ATTGAGACTA	AAGACTGATC	ATGGTTGTAT	GTAAGGAAAA	CATTCTTTTG	AACAGAAATA	1860
	GTGTAATTAA	AAATAATTGA	AAGTGTTAAA	TGTGAACTTG	AGCTGTTTGA	CCAGTCACAT	1920
	TTTTGTATTG	TTACTGTACG	TGTATCTGGG	GCTTCTCCGT	TTGTTAATAC	TTTTTCTGTA	1980
40	TTTGTTGCTG	TATTTTTGGC	ATAACTTTAT	TATAAAAAGC	ATCTCAAATG	CGAAAWAAAA	2040
	АААААААА	AAAAAAC					2058
45							
-	(2) INFORM	ATION FOR SE	EO ID NO: 93	2 •			
		SEQUENCE C	-				
50	1-7	(A) LEN	GTH: 1411 b E: nucleic	ase pairs			
		(C) STR	ANDEDNESS: OLOGY: line	double			
55	(xi) SEQUENCE :			. 92.		
- =				_	AAGCGGAGGA	CTCTCCACCA	60
					TTTCAGATAT		
50				-11000000	**ICHONIAI	contract	120

	CAAGACATCC CCGTAGAAGG AGAAATCACC ATTCCTATGA GATCTCGCAT CCGGGAGTTT	180
	GACAGCTCCA CATTAAATGA ATCTGTTCGC AATACCATCA TGCGTGATCT AAAAGCTGTT	240
5 .	GGGAAAAAT TCATGCATGT TTTGTACCCA AGGAAAAGTA ATACTCTTTT GAGAGATTGG	300
	GATTIGTGGG GCCCTTTGAT CCTTTGTGTG ACACTCGCAT TAATGCTGCA AAGAGACTCT	360
	GCAGATAGTG AAAAAGATGG AGGGCCCCAA TTTGCAGAGG TGTTTGTCAT TGTCTGGTTT	420
10	GGTGCAGTTA CCATCACCCT CAACTCAAAA CTTCTTGGAG GGAACATATC TTTTTTTCAG	480
	AGCCTCTGTG TGCTGGGTTA CTGTATACTT CCCTTGACAG TAGCAATGCT GATTTGCCGG	540
15	CTGGTACTTT TGGCTGATCC AGGACCTGTA AACTTCATGG TTCGGCTTTT TGTGGTGATT	600
	GTGATGTTTG CCTGGTCTAT AGTTGCCTCC ACAGCTTTCC TTGCTGATAG CCAGCCTCCA	660
30	AACCGCAGAG CCCTAGCTGT TTATCCTGTT TTCCTGTTTT ACTTTGTCAT CAGTTGGATG	720
20	ATTCTCACCT TTACTCCTCA GTAAATCAGG AATGGGAAAT TAAAAACCAG TGAATTGAAA	780
	GCACATCTGA AAGATGCAAT TCACCATGGA GCTTTGTCTC TGGCCCTTAT TTGTCTAATT	840
25	TTGGAGGTAT TTGATAACTG AGTAGGTGAG GAGATTAAAA GGGAGCCATA TAGCACTGTC	900
	ACCCCTTATT TGAGGAACTG ATGTTTGAAA GGCTGTTCTT TTCTCTCTTA ATGTCATTTC	960
30	TTTAAAAATA CATGTGCATA CTACACACAG TATATAATGC CTCCTTAAGG CATGATGGAG	1020
30	TCACCGTGGT CCATTTGGGT GACAACCAGT GACTTGGGAA GCACATAGAT ACATCTTACA	1080
	AGTTGAATAG AGTTGATAAC TATTTTCAGT TTTGAGAATA CCAGTTCAGG TGCAGCTCTT	1140
35	AAACACATTG CCTTATGACT ATTAGAATAT GCCTCTCTTT TCATAAATAA AAATACATGG	1200
	TCTATATCCA TTTTCTTTTA TTTCTCTCTC TTAAGCTTAA AAAGGCAATG AGAGAGGTTA	1260
40	GGAGTGGGTT CATACACGGA GAATGAGAAA ACATGCATTA ACCAATATTC AGATTTTGAT	1320
40	CAGGGGAAAT TCTAYACTTG TTGCAAAAAA AAAAAAAAAA AAACTCGAGG GGGGCCCGGT	1380
	ACCCAATCGC NGTATATGAT CGNAAACAAT C	1411
45		
	(2) INFORMATION FOR SEQ ID NO: 93:	
50	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 2187 base pairs (B) TYPE: nucleic acid	
	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
55	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 93:	
	GCTTTGGCTT TTTTTGGCGG ACTGGGGCGC CCTCCGGAAG CGTTTCCAAC TTTCCAGAAG	60
60	TTTCTCGGGA CGGGCAGGAG GGGGTGGGGA CTGCCATATA TAGATCCCGG GAGCAGGGGA	120

	GCGGGCTAAG	AGTAGAATCG	TGTCGCGCTC	GAGAGCGAGA	GTCACGTCCC	GGCGCTAGCC	180
5	CAGCCCGACC	CAGGCCCACC	GTGGTGCACG	CAAACCACTT	CCTGGCCATG	CGCTCCCTCC	240
	TGCTTCTCAG	CGCCTTCTGC	CTCCTGGAGG	CGGCCCTGGC	CGCCGAGGTG	AAGAAACCTG	300
	CAGCCGCAGC	AGCTCCTGGC	ACTGCGGAGA	AGTTGAGCCC	CAAGGCGGCC	ACGCTTGCCG	360
10	AGCGCAGCCG	GCCTGGCCTT	CAGCTTGTAC	CAGGCCATGG	CCAAGGACCA	GGCAGTGGAG	420
	AACATCCTGG	TGTCACCCGT	GGTGGTGGCC	TCGTCGCTGG	GGCTCGTGTC	GCTGGGCGGC	480
15	AAGGCGACCA	CGGCGTCGCA	GGCCAAGGCA	GTGCTGAGCG	CCGAGCAGCT	GCGCGACGAG	540
	GAGGTGCACG	CCGGCCTGGG	CGAGCTGCTG	CGCTCACTCA	GCAACTCCAC	GGCGCGCAAC	600
	GTGACCTGGA	AGCTGGGCAG	CCGACTGTAC	GGACCCAGCT	CAGTGAGCTT	CGCTGATGAC	660
20	TTCGTGCGCA	GCAGCAAGCA	GCACTACAAC	TGCGAGCACT	CCAAGATCAA	CTTCCGCGAC	720
	AAGCGCAGCG	CGCTGCAGTC	CATCAACGAG	TGGGCCGCGC	AGACCACCGA	CGGCAAGCTG	780
25	CCCGAGGTCA	CCAAGGACGT	GGAGCGCACG	GACGGCGCCC	TGTTAGTCAA	CGCCATGTTC	840
	TTCAAGCCAC	ACTGGGATGA	GAAATTCCAC	CACAAGATGG	TGGACAACCG	TGGCTTCATG	900
	GTGACTCGGT	CCTATACCGT	GGGTGTCATG	ATGATGCACC	GGACAGGCCT	CTACAACTAC	960
30	TACGACGACG	AGAAGGAAAA	GCTGCAAATC	GTGGAGATGC	CCCTGGCCCA	CAAGCTCTCC	1020
	AGCCTCATCA	TCCTCATGCC	CCATCACGTG	GAGCCTCTCG	AGCGCCTTGA	AAAGCTGCTA	1080
35	ACCAAAGAGC	AGCTGAAGAT	CTGGATGGGG	AAGATGCAGA	AGAAGGCTGT	TGCCATCTCC	1140
	TTGCCCAAGG	GTGTGGTGGA	GGTGACCCAT	GACCTGCAGA	AACACCTGGC	TGGGCTGGGC	1200
	CTGACTGAGG	CCATTGACAA	GAACAAGGCC	GACTTGTCAC	GCATGTCAGG	CAAGAAGGAC	1260
40	CTGTACCTGG	CCAGCGTGTT	CCACGCCACC	GCCTTTGAGT	TGGACACAGA	TGGCAACCCT	1320
	TTGACCAGAA	TTACGGGCGG	AGGAGTGCGC	ACCCAAGTGT	TCTACGCCGA	CCACCCCTTC	1380
45	ATTTCCTAGT	GCGGGACACC	CAAAGCGGTC	CCTGCTATTC	ATTGGGCGCC	TGGTCCGGCC	1440
	TAAGGGTGAC	AAGATGCGAG	ACGAGTTATA	GGCCTCAGGG	TGCACACAGG	ATGGCAGGAG	1500
	GCATCCAAAG	GCTCCTGAGA	CACATGGGTG	CTATTGGGGT	TGGGGGGGAG	GTGAGGTACC	1560
50	AGCCTTGGAT	ACTCCATGGG	GTGGGGTGGA	AAAGCAGACC	GGGGTTCCCG	TGTGCCTGAG	1620
	CGGACTTCCC	AGCTAGAATT	CACTCCACTT	GGACATGGGC	CCCAGATACC	ATGATGCTGA	1680
55	GCCCGGAAAC	TCCACATCCT	GTGGGACCTG	GGCCATAGTC	ATTCTGCCTG	CCCTGAAAGT	1740
_	CCCAGATCAA	GCCTGCCTCA	ATCAGTATTC	ATATTTATAG	CCAGGTACCT	TCTCACCTGT	1800
	GAGACCAAAT	TGAGCTAGGG	GGGTCAGCCA	GCCCTCTTCT	GACACTAAAA	CACCTCAGCT	1860
60	GCCTCCCCAG	CTCTATCCCA	ACCTCTCCCA	ACTATAAAAC	TAGGTGCTGC	AGCCCCTGGG	1920

	ACCAGGCACC CCCAGAATGA CCTGGCCGCA GTGAGGCGGA TTGAGAAGGA GCTCCCAGGA	1960
_	GGGGCTTCTG GGCAGACTCT GGTCAAGAAG CATCGTGTCT GGCGTTGTGG GGATGAACTT	2040
5	TTTGTTTTGT TTCTTCCTTT TTTAGTTCTT CAAAGATAGG GAGGGAAGGG GGAACATGAG	2100
	CCTTTGTTGC TATCAATCCA AGAACTTATT TGTACATTTT TTTTTTCAAT AAAACTTTTC	2160
10	CAATGACAAA AAAAAAAAA AAAAAAA	2187
15	(2) INFORMATION FOR SEQ ID NO: 94:	
	(i) SEQUENCE CHARACTERISTICS:	
20	(A) LENGTH: 757 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double	
20	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 94:	
25	GACAGTACGG TCGGATTCCC GGGTCGACCC ACGCGTCCGC GGACGGTGAA GAAGGTGAAG	60
	ATGGCGGTGG CCAGGGCCGG GGTCTTGGGA GTCCAGTGGC TGCAAAGGGC ATCCCGGAAC	120
30	GTGATGCCGC TGGGCGCACG GACAGCCTCC CACATGACCA AGGACATGTT CCCGGGGCCC	180
30	TATCCTAGGA CCCCAGAAGA ACGGGCCGCC GCCGCCAAGA AGTATAATAT GCGTGTGGAA	240
	GACTACGAAC CTTACCCGGA TGATGGCATG GGGTATGGCG ACTACCCGAA GCTCCCTGAC	300
35	CGCTCACAGC ATGAGAGAGA TCCATGGTAT AGCTGGGACC AGCCGGGCCT GAGGTTGAAC	360
	TGGGGTGAAC CGATGCACTG GCACCTAGAC ATGTACAACA GGAACCGTGT GGATACATCC	420
40	CCCACACCTG TTTCTTGGCA TGTCATGTGT ATGCAGCTCT TCGGTTTCCT GGCTTTCATG	480
	ATATTCATGT GCTGGGTGGG GGACGTGTAC CCTGTCTACC AGCCTGTGGG ACCAAAGCAG	540
	TATCCTTACA ATAATCTGTA CCTGGAACGA GGCGGTGATC CCTCCAAAGA ACCAGAGCGG	600
45	GTGGTTCACT ATGAGATCTG AGGAGGCTTC GTGGGCTTTT GGGTCCTCTA ACTAGGACTC	660
	CCTCATTCCT AGAAATTTAA CCTTAATGAA ATCCCTAATA AAACTCAGTG CTGTGTTAAA	720
50	AAAAAAAAA AAAAAAAA AAAAAGGGGG GCCCCNN	757
	(2) INFORMATION FOR SEQ ID NO: 95:	
55	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 2394 base pairs (B) TYPE: nucleic acid	
60	(C) STRANDEDNESS: double(D) TOPOLOGY: linear	

(xi)	SEQUENCE	DESCRIPTION:	SEQ	ID	NO:	95:
------	----------	--------------	-----	----	-----	-----

5	GGCACGAGCA	CTCCTGCACT	TCCCCACCCC	CACGACCGAA	CCTGGCTTCG	CTAACGCCCT	60
5	CCCAGCTCCC	TCGGGCCTGA	CTTCCGGTTT	CCTCGCGCGT	CCCTGGCGCC	GAGCCGCGGA	120
	CAGCAGCCCC	TTTTCCGGCT	GAGAGCTCAT	CCACACTTCC	AATCACTTTC	CGGAGTGCTT	180
10	CCCCTCCCTC	CGGCCCGTGC	TGGTCCCGAC	GCCGCCCTG	GGTCTCGCGC	GCGTATTGCT	240
	GGGTAACGGG	CCTTCTCYCG	CGTCGGCCCG	GCCCCTCCTG	CCTCGGCTCG	TCCCTCCTTC	300
15	CAGAACGTCC	CGGGCTCCTG	CCGAGTCAGA	AGAAATGGGA	CTCCCTCCGC	GACGTGCCCG	360
	GAGCAGCTCC	CTTCGCTGTG	GAAGCGGCGG	TGTCTTCGAA	GAAACCGGAA	GCCCGTGGTG	420
	ACCCCTGGCG	ACCCGGTTTG	TTTTCGGTCC	GTTTCCAAAC	ACTAAGGAAT	CGAAACTCGG	480
20	CGGCCTTGGG	GGCGGCCCTA	CGTAGCCTGG	CTTCTGGTTG	TCATGGATGC	ACTGGTAGAA	540
	GATGATATCT	GTATTCTGAA	TCATGAAAAA	GCCCATAAGA	GAGATACAGT	GACTCCAGTT	600
25	TCAATATATT	CAGGAGATGA	ATCTGTTGCT	TCCCATTTTG	CTCTTGTCAC	TGCATATGAA	660
	GACATCAAAA	AACGACTTAA	GGATTCAGAG	AAAGAGAACT	CTTTGTTAAA	GAAGAGAATA	720
	AGATTTTTGG	AAGAAAAGCT	AATAGCTCGA	TTTGAAGAAG	AAACAAGTTC	CGTGGGACGA	780
30	GAACAAGTAA	ATAAGGCCTA	TCATGCATAT	CGAGAGGTTT	GCATTGATAG	AGATAATTTG	840
	AAGAGCAAAC	TGGACAAAAT	GAATAAAGAC	AACTCTGAAT	CTTTGAAAGT	ATTGAATGAG	900
35	CAGCTACAAT	CTAAAGAAGT	AGAACTCCTC	CAGCTGAGGA	CAGAGGTGGA	AACTCAGCAG	960
	GTGATGAGGA	ATTTAAATCC	ACCTTCATCA	AACTGGGAGG	TGGAAAAGTT	GAGCTGTGAC	1020
	CTGAAGATCC	ATGGTTTGGA	ACAAGAGCTG	GAACTGATGA	GGAAAGAATG	TAGCGATCTC	1080
40	AAAATAGAAC	TACAGAAAGC	CAAACAAACG	GATCCATATC	AGGAAGACAA	TCTGAAGAGC	1140
	AGAGATCTCC	AAAAACTAAG	CATTTCAAGT	GATAATATGC	AGCATGCATA	CTGGGAACTG	1200
45	AAGAGAGAAA	TGTCTAATTT	ACATCTGGTG	ACTCAAGTAC	AAGCTGAACT	ACTAAGAAAA	1260
	CTGAAAACCT	CAACTGCAAT	CAAGAAAGCC	TGTGCCCCTG	TAGGATGCAG	TGAAGACCTT	1320
	GGAAGAGACA	GCACAAAACT	GCACTTGATG	AATTTTACTG	CAACATACAC	AAGACATCCC	1380
50	CCTCTCTTAC	CAAATGGCAA	AGCTCTTTGT	CATACCACAT	CTTCCCCTTT	ACCAGGAGAT	1440
	GTAAAGGTTT	TATCAGAGAA	AGCAATCCTC	CAATCATGGA	CAGACAATGA	GAGATCCATT	1500
55	CCTAATGATG	GTACATGCTT	TCAGGAACAC	AGTTCTTATG	GCAGAAATTC	TCTGGAAGAC	1560
	AATTCCTGGG	TATTTCCAAG	TCCTCCTAAA	TCAAGTGAGA	CAGCATTTGG	GGAAACTAAA	1620
	ACTAAAACTT	TGCCTTTACC	CAACCTTCCA	CCACTGCATT	ACTTGGATCA	ACATAATCAG	1680
60	AACTGCCTTT	ATAAGAATTA	ATTTGGAAGA	GATTCACGAT	TTCACCATGA	GGACACTTAT	1740

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	CTCTTTCAGT GGTCCTCCCA AGAAATTATT TAACAAACTG AANGGAGATT TTGATTAAAA	1800
5	TTTTGCAGAG GTCTTCAGTA TCTATATTTG AACACACTGT ACAATAGTAC AAAAACCAAC	1860
5	ATAGTTGGTT TTCTAGTATG AAAGAGCACC CTCTAGCTCC ATATTCTAAG AATCTGAAAT	1920
	ATGCTACTAT ACTAATTAAT AAGTAAACTT AAGGTGTTTA AAAAACTCTG CCTTCTATAT	1980
10	TAATTGTAAA ATTTTGCCTC TCAGAAGAAT GGAATTGGAG ATTGTAGACG TGGTTTTACA	2040
	AAATGTGAAA TGTCTAAATA TCTGTTCATA AAAATAAAAG GAAAACATGT TTCTTCAAAT	2100
1.5	TGCATAATGG AACAAATGGC AATGTGAGTA GGTTACATTT CTGTTGTTAT AATGCGTAAA	2160
15	GATATTGAAA ATATAATGAA ATAAAAGCAT CTTAGGTTAT ACCATCTTTA TATGCTATTG	2220
	CGTTTCAATA TTTAAGATTT AAAGTGATTT TTTGGTCACA GTGTTTTGTT GATAAAATTT	2280
20	TTTTAGAATT GAAGTTTGAA TTCTAAGACT TGAAACAACC TGATCACTGA AGCCAACTTT	2340
	GTCCCAGCAC ATTCCTTAAG TCCTAATTGG GGAAAAAAAA AAAAAAAAAC TCGA	2394
25		
23	(2) INFORMATION FOR SEQ ID NO: 96:	
	(i) SEQUENCE CHARACTERISTICS:	
30	(A) LENGTH: 672 base pairs	
	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
25	(D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NC: 96:	
35	AGTGCTCTGT TGCCCAGGCT GGAGTGCGTT AGTGTAATGT CAGTCCACTG CAACCTCCAC	60
		120
40	CCCCAGGTTC AAGCAATTCT CATGCCTCAG CCTCCCAAGT AGCTGAAATT ACTGGCATGC	180
	ACCACCACAC CCAGCTGATG TTTATTTATT TATTTATATA TTTATTTATT TTAGGTGTTT	
	TTTTTTTTT TTTTTGAGAC GGAGTCTTGC TCTGTTGCCC TGGGTGTGT TACGTGGRAT	240
45	TACCATYCTG GGTGACTCAC TGAAATGTAC TCMCAGTGAG TCATGCCTTC MAATGACATC	300
	TCAAGTTCTG CCTGCTTGGA GATACATCTG GGGATCTTAA GGGGTGAGGG ACTACTCAAC	
50	AAGAAGGAAT TTAGCCTGTC TTTTTAAATA AACGGCATTT CTTTTTCCTA KAAAAATGGG	420
	AAATTCTTCA ATTCTCTAAT ACAGGGACAC TGAGATAACA AAGAGGAAAG TGTCTGGTTG	480
	GAGGTTGGGA RGCCACCCTG GGGTCTCTCC TACAAAAATG GAAAAGAAAA	540
55	AAATCMAGCA AAGCACAARA AAKTTTCCCT TTGCTAAAAG GGAAAAGATG CCCCMCAATG	600

CCCATAAACA TGAACTGGGG ATAAGGAGGA RAATGTCTCT YCTTGGCACC CCCAAACAAA

660

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60

CGTTAATTAC CC

5	(2) INFORM	ATION FOR S	EQ ID NO: 9	7:			
J	(i)	(B) TYP	HARACTERIST GTH: 1419 b E: nucleic ANDEDNESS:	ase pairs acid			
10			OLOGY: line				
	(xi) SEQUENCE	DESCRIPTION	: SEQ ID NO	: 97:		
15	TAAGAACAGA	ACAGCAAGTA	TGAACCACAT	GGAACTTAAA	ACATATGGGT	GTGAAGTCCA	60
13	CTTATGTAGA	CAAAACTTAT	AATTTCCAAA	CTGTTGTCTA	GTATACAGTG	ATCAGTTGCT	120
	CTCTGTTCAA	GTCATTCCAC	ACATTTCCCT	ATTTTAGGCT	ATTATAATAT	AGAAAGAAAA	180
20	TGGGAAGCAT	TAGTTGGAGC	TAGAAAATGA	ACTGTATATT	ATTGCTATAT	TTGCTAATAC	240
	CAACTATTTC	AATAAGTGTT	GTACCATATG	TAGCATTAAA	ТАТААААТАС	ATAAAAGAAT	300
25	GTACAGAAAA	TAGCTTTTAT	TGAGTAATAT	TACATTTCAT	TTATACTGTA	GCAATATATT	360
23	TGTAGGTATA	CTCTGTAAGG	GCTTTAAATA	AAAGAGGTCC	ATTAATACTT	ССТТАТАААА	420
	ATTCTAGTCT	GTTTCATTAC	TGCCCAGATG	TTTTAGAGAT	AAATATTTAT	GCAGAAGGTA	480
30	TTTTKGAAAG	TCYCCYTTTG	TCTGATAGAG	TTTAACNAGA	TATTTAAATT	TAGTGCYCNA	540
	GAAATCCCAC	AAGTCACGGT	CTAAACACAC	TTAGAATACT	ACAGCATAAA	TCTGTTAGCA	600
35	TTANTTGCCA	AATAAGACAG	TTGGGATCCC	AAACCCCAAG	TCCTTGAGCA	ATGTTTTTCC	660
55	TCAAAAAGCT	GCTATNCCAA	TGATATAGGA	AAAWACATTG	TGTTTTCCTA	AACACACTTT	720
	TCTTTTTAAA	TGTGCTTCAT	TGTTTGATTT	GGTCCTGCCT	AAATTTCACA	AGCTAGGCCA	780
40	ATGAAGGCTG	AATCAAAGAC	ATTTCATCCA	CCAATATCAT	GTGTAGATAT	TATGTATAGA	840
	AAATAAAATA	AATTATGGCT	CTAACTTCTG	TGTTGCTGTT	TATCTTGTTA	TTTTTCGGCG	900
45	TTATACTAAT	GNGTTTATTG	AGAGCATTTT	ACCTTCCAGA	CTTCTCATGG	CTAACTTTTG	960
15	GTCTGWATTT	TGSTCCTTAG	ATGKGAATAT	TTCTTATTAG	TYTGCTYCCT	GCWACGCAAT	1020
	GACTGCATTT	CTATCATTTC	TCAGTTTGTT	AGWATATGTG	GATAGTATTC	TACTGTATAA	1080
50	ATGATTGCAA	AGTTTATCAA	AAACAAATTA	TTATATGTAG	CTTTTCTACA	GTGCTTTGCT	1140
	AAACCATGTA	GTACTAGTTA	AGTSTTCCTT	GAAAATAAAG	ATACACTCTT	ATAGGGGACA	1200
55	GTTCCTGTTC	ACTCCCAGGA	AACTTTTTTA	AAAGATGACA	CTGAATGTTT	ATTGCACTTT	1260
55	AGTGCAGTGA	AGTGGCAATA	AAACCTAACA	TGAATCAAGG	TTGTTTATGG	CAGATGCATG	1320
	TGTTGCTTTA	CAGAGTTTAG	CAAAAGCTCT	TAATTTTATG	TCATACTGTA	TTCTACTGAA	1380
60	TAATAAAGCT	AACATTATTC	ААТААТААА	TGGAAAAA			1419

5 (2) INFORMATION FOR SEQ ID NO: 98:

10

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1487 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 98:

	(X1) SEQUENCE DESCRIPTION: SEQ ID NO. 96.	
15	GCGACCGCGC CCCTTTCAGC TAGCTCGCTC GCTCGCTCTG CTTCCCTGCT GCCGGCTGCG	60
	CATGGCKWTG GCGTTGGCGG CGCTGGCGGC GGCTGCGCAG CCGGTACCAG	120
20	CAGTTGCAGA ATGAAGAAGA GTCTGGAGAA CCTGAACAGG CTGCAGGTGA TGCTCCTCCA	180
20	CCTTACAGCA GCATTTCTGC AGAGAGCGCA GTTTTCCACC TATTTCCCTG GATATTTTGA	240
	TGGTCAGTAC TGGCTCTGGT GGGTGTTCCT TGTTTTAGGC TTTCTCCTGT TTCTCAGAGG	300
25	ATTTATCAAT TATGCAAAAG TTCGGAAGAT GCCAGAAACT TTCTCAAATC TCCCCAGGAC	360
	CAGAGTTCTC TTTATTTATT AAAGATGTTT TCTGGCAAAG GCCTTCCTGC ATTTATGAAT	420
20	TCTCTCTCAA GAAGCAAGAG AACACCTGCA GGAAGTGAAT CAAGATGCAG AACACAGAGG	480
30	AATAATCACC TGCTTTAAAA AAATAAAGTA CTGTTGAAAA GATCATTTCT CTCTATTTGT	540
	TCCTAGGTGT AAAATTTTAA TAGTTAATGC AGAATTCTGT AATCATTGAA TCATTAGTGG	600
35	TTAATGTTTG AAAAAGCTCT TGCAATCAAG TCTGTGATGT ATTAATAATG CCTTATATAT	660
	TGTTTGTAGT CATTTTAAGT AGCATGAGCC ATGTCCCTGT AGTCGGTAGG GGGCAGTCTT	720
40	GCTTTATTCA TCCTCCATCT CAAAATGAAC TTGGAATTAA ATATTGTAAG ATATGTATAA	780
40	TGCTGGCCAT TTTAAAGGGG TTTTCTCAAA AGTTAAACTT TTGTTATGAC TGTGTTTTTG	840
	CACATAATCC ATATTTGCTG TTCAAGTTAA TCTAGAAATT TATTCAATTC TGTATGAACA	900
45	CCTGGAAGCA AAATCATAGT GCAAAAATAC ATTTAAGGTG TGGTCAAAAA TAAGTCTFTA	960
	ATTGGTAAAT AATAAGCATT AATTTTTTAT AGCCTGTATT CACAATTCTG CGGTACCTTA	1020
50	TTGTACCTAA GGGATTCTAA AGGTGTTGTC ACTGTATAAA ACAGAAAGCA CTAGGATACA	1080
50	AATGAAGCTT AATTACTAAA ATGTAATTCT TGACACTCTT TCTATAATTA GCGTTCTTCA	1140
	CCCCCACCC CACCCCCACC CCCCTTATTT TCCTTTTGTC TCCTGGTGAT TAGGCCAAAG	1200
55	TCTGGGAGTA AGGAGAGGAT TAGGTACTTA GGAGCAAAGA AAGAAGTAGC TTGGAACTTT	1260
	TGAGATGATC CCTAACATAC TGTACTACTT GCTTTTACAA TGTGTTAGCA GAAACCAGTG	1320
	GGTTATAATG TAGAATGATG TGCTTTCTGC CCAAGTGGTA ATTCATCTTG GTTTGCTATG	1380
60		

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	TTAAAACTGT	AAATACAACA	GAACATTAAT	AAATATCTCT	TGTGTAGCAC	CTTTAAAAAA	1440
	AAAAAAAAA	AAAAAAAA	AAAAAAAAN	ccceeeee	GGCCCCN		1487
5							
		ATION FOR SE					
10	(i)	(B) TYP (C) STR	HARACTERIST GTH: 1653 b E: nucleic ANDEDNESS: OLOGY: line	ase pairs acid double			
15	(xi) SEQUENCE I	DESCRIPTION	: SEQ ID NO	: 99:		
	GCGACCGCGC	CCTTCAGCTA	GCTCGCTCGC	TCGCTCTGCT	TCCCTGCTGC	CGGCTGCGCA	60
20	TGGCTTNGGC	GTTGGCGGCG	CTGGCGGCGG	CTCGAGCCGC	CTGCGSAGCC	GGTACCAGCA	120
	GTTGCAGAAT	GAAGAAGAGT	CTGGAGAACC	TGAACAGGCT	GCAGGTGATG	CTCCTCCACC	180
25	TTACAGCAGC	ATTTCTGCAG	AGAGCGCACA	TNATTTTGAC	TACAAGGATG	AGTCTGGGTT	240
25	TCCAAAGCCC	CCATCTTACA	ATGTAGCTAC	AACACTGCCC	AGTTATGATG	AAGCGGAGAG	300
	GACCAAGGCT	GAAGCTACTA	TCCCTTTGGT	TCCTGGGAGA	GATGAGGATT	TTGTGGGTCG	360
30	GGATGATTTT	GATGATGCTG	ACCAGCTGAG	GATAGGAAAT	GATGGGATTT	TCATGTTAAC	420
	TTTTTCATG	GCATTCCTCT	TTAACTGGAT	TGGGTTTTTC	CTGTCTTTT	GCCTGACCAC	480
35	TTCAGCTGCA	GGAAGGTATG	GGGCCATTTC	AGGATTTGGT	CTCTCTCTAA	TTAAATGGAT	540
55	CCTGATTGTC	AGGTTTTCCA	CCTATTTCCC	TGCATTTATG	AATTCTCTCT	CAAGAAGCAA	600
	GAGAACACCT	GCAGGAAGTG	AATCAAGATG	CAGAACACAG	AGGAATAATC	ACCTGCTTTA	660
40	AAAAATAAA	GTACTGTTGA	AAAGATCATT	TCTCTCTATT	TGTTCCTAGG	TGTAAAATTT	720
	TAATAGTTAA	TGCAGAATTC	TGTAATCATT	GAATCATTAG	TGGTTAATGT	TTGAAAAAGC	780
45	TCTTGCAATC	AAGTCTGTGA	TGTATTAATA	ATGCCTTATA	TATTGTTTGT	AGTCATTTTA	840
	AGTAGCATGA	GCCATGTCCC	TGTAGTCGGT	AGGGGGCAGT	CTTGCTTTAT	TCATCCTCCA	900
	TCTCAAAATG	AACTTGGAAT	TAAATATTGT	AAGATATGTA	TAATGCTGGC	CATTTTAAAG	960
50	GGGTTTTCTC	AAAAGTTAAA	CTTTTGTTAT	GACTGTGTTT	TTGCACATAA	TCCATATTTG	1020
	CTGTTCAAGT	TAATCTAGAA	ATTTATTCAA	TTCTGTATGA	ACACCTGGAA	GCAAAATCAT	1080
55	AGTGCAAAAA	TACATTTAAG	GTGTGGTCAA	AAATAAGTCT	TTAATTGGTA	AATAATAAGC	1140
	ATTAATTTT	TATAGCCTGT	ATTCACAATT	CTGCGGTACC	TTATTGTACC	TAAGGGATTC	1200
	TAAAGGTGTT	GTCACTGTAT	AAAACAGAAA	GCACTAGGAT	ACAAATGAAG	CTTAATTACT	1260
60	AAAATGTAAT	TCTTGACACT	CTTTCTATAA	TTAGCGTTCT	TCACCCCCAC	CCCCACCCC	1320

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	ACCCCCCTTA TTTTCCTTTT GTCTCCTGGT GATTAGGCCA AAGTCTGGGA GTAAGGAGAG	1380
_	GATTAGGTAC TTAGGAGCAA AGAAAGAAGT AGCTTGGAAC TTTTGAGATG ATCCCTAACA	1440
5	TACTGTACTA CTTGCTTTTA CAATGTGTTA GCAGAAACCA GTGGGTTATA ATGTAGAATG	1500
	ATGTGCTTTC TGCCCAAGTG GTAATTCATC TTGGTTTGCT ATGTTAAAAC TGTAAATACA	1560
10	ACAGAACATT AATAAATATC TCTTGTGTAG CACCTTTTAW AAAAAAAAAA AAAAAAAAAA	1620
	AAAAAAAAA AAAAANCCCG GGGGGGGCC CCN	1653
15		
	(2) INFORMATION FOR SEQ ID NO: 100:	
20	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1145 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
25	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 100:	
	TTTTTTTTT TTTTTTTTT TTGACTGAAC TAAGTGGCTT TTTTATTAGA GAAAGCCAGA	60
30	ATTACAAAAG ACTTCCCTTT TCTTGGGGTA TGGCTGTCTC AGCACAATAC TCAACATAAC	120
30	TGCAGAACTG ATGTGGCTCA GGCACCCTGG TTTTAATTCC TTGAGGATCT GGCAATTGGC	180
	TTACGCAAAA GGTCACCATT TGAGGTCCTG CCTTACTAAT TATGTGCTGC CCAACAACTA	240
35	AATTTGTAAT TTGTTTTCT CTAGTTTGAG CAGGGTCTGA ATTTTTTCAT TTATTTCCTT	300
	TTTTGCCAGC AGACAGACTT GAGTCTGTAA AGACAAGCAA ATACACTGAC AGAAGTTTAC	360
40	CATAGTTTCT AAAATGTAAA AAAGAAAACC CCCAAAAGAC TCAAGAAAAT TAGACCACAA	420
40	ATTITGCATT GTTCATTGTA GCACTATTGG TAATAAAATA ACAAATGTTT GTGCATTTTT	480
	ATGTGAAGAT CCTTCTCGTA TTTCATTTGG AAAGATGAGC AAGAGGTCTG CTTCCTTCAT	540
45	TTTACTTCCC CTTCTGTTTT TGAAAGGCAG TTTCGCCAAG CTTAATGCAA GAATATCTGA	600
	CTGTTTAGAA GAAAGATATT GCCACAATCT CTGGATGGTT TTCCAGGGTT GTGTTATTAC	660
50	TGAGCTTCAT CTTTCCAGAA TGAGCAAAAC ACTGTCCAGT CTTTGTTACG ATTTTGTAAT	720
50	AAATGTGTAC ATTTTTTTTA AATTTTTGGA CATCACATGA ATAAAGGTAT GTATGTACGA	780
	ATGTGTATAT ATTATATATA TGACATCTAT TTTGGAAAAT GTTTGCCCTG CTGTACCTCA	840
55	TTTTTAGGAG GTGTGCATGG ATGCAATATA TGAAAATGGG ACATTCTGGA ACTGCTGGTC	900
	AGGGGACTTT GTCGCCCTGT GCACTAAAAG GGCCAGATTT TCAGCAGCCA AGGACATCCA	960

TACCCAAGTG AATGTGATGG GACTTAAAAG AAGTGAACTG AGACAATTCA CTCTGGCTGT

1020

	TTGAACAGCA GCGTTTCATA GGAAGAGAAA AAAAGATCAA TCTTGTATTT TCTGACCACA	1080
	TAAAGGCTTC TTCTCTTTGT AATAAAGTAG AAAAGCTCTC CTCAAAAAAA AAAAAAAAA	1140
5	· AAAAA	1145
10	(2) INFORMATION FOR SEQ ID NO: 101:	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 734 base pairs	
1.5	(B) TYPE: nucleic acid	
15	(C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 101:	
20	TACCCGGCGG ATTCCAGGAA GGTAAATTTA GTCCTATAAT TTTCAGCTTA ATTATAAACA	60
	AAGGAACAAA TAAGTGGAAG GGCAGCTATT ACCATTCGCT TAGTCAAAAC ATTCGGTTAC	120
25	TGCCCTTTAA TACACTCCTA TCATCAGCAC TTCCACCATG TATTACAAGT CTTGACCCAT	180
	CCCTGTCGTA ACTCCAGTAA AAGTTACTGT TACTAGAAAA TTTTTATCAA TTAACTGACA	240
	AATAGTTTCT TTTTAAAGTA GTTTCTTCCA TCTTTATTCT GACTAGCTTC CAAAATGTGT	300
30	TCCCTTTTTG AATCGAGGTT TTTTTGTTTT GTTTTGTTTT	360
	TGTGCTTCTA TTGCTTTTTT GTGTTTTGTT AAGCATGTCC CTTGGCCCAA ATGGAAGAGG	420
35	AAATGTTTAA TTAATGCTTT TTAGTTTAAA TAAATTGAAT CATTTATAAT AATCAGTGTT	480
	AACAATTTAG TGACCCTTGG TAGGTTAAAG GTTGCATTAT TTATACTTGA GATTTTTTTC	540
	CCCTAACTAT TCTGTTTTTT GTACTTTAAA ACTATGGGGG AAATATCACT GGTCTGTCAA	600
40	GAAACAGCAG TAATTATTAC TGAGTTAAAT TGAAAAGTCC AGTGGACCAG GCATTTCTTA	660
	TATAAATAAA ATTGGTGGTA CTAATGTGAA AAAAAAAAAA	720
45	CCGGTACCCT ATTA	734
	(2) INFORMATION FOR SEQ ID NO: 102:	
50	(i) SPONENCE CHARACTERISTICS.	
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 713 base pairs	
	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
55	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 102:	
60	CCGCGGGAAC GCTGTCCTGG CTGCCGNCAC CCGAACAGCC TGTCCTGGTG CCCCGGCTCC	60

253

	CTGCCCCGCG CCCAGTCATG ACCCTGCGCC CCTCACTCCT CCCGCTCCAT CTGCTGCTGC	120					
	TGCTGCTGCT CAGTGCGGCG GTGTGCCGGG CTGAGGCTGG GCTCGAAACC GAAAGTCCCG	180					
5	TCCGGACCCT CCAAGTGGAG ACCCTGGTGG AGCCCCCAGA ACCATGTGCC GAGCCCGCTG	240					
	CTTTTGGAGA CACGCTTCAC ATACACTACA CGGGAAGCTT GGTAGATGGA CGTATTATTG	300					
10	ACACCTCCCT GACCAGAGAC CCTCTGGTTA TAGAACTTGG CCAAAAGCAG GTGATTCCAG	360					
10	GTCTGGAGCA GAGTCTTCTC GACATGTGTG TGGGAGAGAA GCGAAGGGCA ATCATTCCTT	420					
	CTCACTTGGC CTATGGAAAA CGGGGATTTC CACCATCTGT CCCAGCGGAT GCAGTGGTGC	480					
15	AGTATGACGT GGAGCTGATT GCACTAATCC GAGCCAACTA CTGGCTAAAG CTGGTGAAGG	540					
	GCATTTTGCC TCTGGTAGGG ATGGCCATGG TGCCACCCTC CTGGGCCTCA TTGGGTATCA	600					
20	CCTATACAGA AAGGCCAATA GACCCAAAAGT CTCCAAAAAG AAGCTCAAGG AAGAGAAACG	660					
20	AAACAAGAGC AAAAAGAAAT AATAAATAAT AAATTTTAAA AAACTTAAAA AAA	713					
25	(2) INFORMATION FOR SEQ ID NO: 103:						
30	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1080 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear						
35	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 103:						
33	CCGATGTGGA CATCATCCTG TCTATCCCCA TGTTCCTGCG CCTGTACCTG ATCGCCCGAG	60					
	TCATGCTGCT GCACAGAAGC TCTTCACCGA TGCCTCGTCC CGCAGCATCG GGGCCCTCAA	120					
40	CAAGATCAAC TTCAACACCC GCTTTGTCAT GAAGACGCTC ATGACCATCT GCCCTGGCAC	180					
	TGTGCTGCTC GTGTTCAGCA TCTCTCTGTG GATCATTGCT GCCTGGACCG TCCGTGTCTG	240					

TGAAAGTCCT GAATCACCAG CCCAGCCTTC TGGCTCATCA CTTCCTGCTT GGTACCATGA 300 45 CCAGCAGGAC GTAACTAGTA ACTTTCTGGG TGCCATGTGG CTCATCTCCA TCACATTCCT 360 420 TTCCATTGGT TATGGGGACA TGGTGCCCCA CACATACTGT GGGAAAGGTG TCTGTCTCCT CACTGGCATC ATGGGTGCAG GCTGCACTGC CCTTGTGGTG GCCGTGGTGG CCCGAAAGCT 480 50 GGAACTCACC AAAGCGGAGA AGCACGTTCA TAANTTCATG ATGGACACTC AGCTCACCAA 540 GCGGATCAAG AATGYTGCAG CCAATGTCCT TSGGGAAACA TGGTTAATCT ATAAACACAC 600 55 AAAGYTGYTA AAGAAGATTG ACCATGCCAA AGTGAGGAAC ACCAGAGGAA GTTCYTCCAA 660 720 GTATCCACCA GTTGAGGAGC GTCAAGATGG AACAGAGGAA GCTGAGTGAC CAAGCCAACA NTCTGGTGGA CCTTTCCAAG ATGCAGAATG TCMTGTATGA CTTAATCACA GAACTCAATG 780 60

	ACCGGAGCGA AGACCTGGAG AAGCAGATTG GCAGCCTGGA GTCGAAGCTG GAGCATCTCA	840
5	CCGCCAGCTT CAACTCCCTG CCGCTGCTCA TCGCCGACAC CCTGCGCCAG CAGCAGCAGC	900
,	AGCTCCTGTC TGCCATCATC GAGGCCCGGG GTGTCAGCGT GGCAGTGGGC ACCACCCACA	960
	CCCCAATCTC CGATAGCCCC ATTGGGGTCA GCTCCACCTC CTTCCCGACC CCGTACACAA	1020
10	GTTCAAGCAG TTGCTAAATA AATCTCCCCA CTCCAGAAGC ATTAAAAAAA AAAAAAAAAA	1080
15	(2) INFORMATION FOR SEQ ID NO: 104:	
20	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 489 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 104:	
25	GGCACGAGAG GCTTTGAAGC ATTTTTGTCT GTGCTCCCTG ATCTTCAGGT CACCACCATG	60
	AAGTTCTTAG CAGTCCTGGT ACTCTTGGGA GTTTCCATCT TTCTGGTCTC TGCCCAGAAT	120
30	CCGACAACAG CTGCTCCAGC TGACACGTAT CCAGCTACTG GTCCTGCTGA TGATGAAGCC	180
50	CCTGATGCTG AAACCACTGC TGCTGCAACC ACTGCGACCA CTGCTGCTCC TACCACTGCA	240
	ACCACCGCTG CTTCTACCAC TGCTCGTAAA GACATTCCAG TTTTACCCAA ATGGGTTGGG	300
35	GATCTCCCGA ATGGTAGAGT GTGTCCCTGA GATGGAATCA GCTTGAGTCT TCTGCAATTG	360
	GTCACAACTA TTCATGCTTC CTGTGATTTC ATCCAACTAC TTACCTTGCC TACGATATCC	420
40	CCTTTATCTC TAATCAGTTT ATTTTCTTTC AAATAAAAA TAACTATGAG CAACAAAAA	480
	ААААААА	489
45	(2) INFORMATION FOR SEQ ID NO: 105:	
50	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 640 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
55	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 105:	
55	GCGGTCGCCG CTGTTGTTGT GGTCCCCATG GAGCTGCCGT AGCGGACCCA GCACAGCCAG	60
	GAGCGTCCGG GATGAGCTCA GCCGCGGCCG ACCACTGGGC GTGGTTGCTG GTGCTCAGCT	120
60	TCGTGTTTGG ATGCAATGTT CTTAGGATCC TCCTCCCGTC CTTCTCATCC TTCATCTCA	180

	GGGTGCTGCA	GAAGGACGCG	GAGCAGGAGT	CACAGATGAG	AGCGGAGATC	CAGGACATGA	240
_	AGCAGGAGCT	CTCCACAGTC	AACATGATGG	ACGAGTTTGC	CAGATATGCC	AGGCTGGAAA	300
3	GAAAGATCAA	CAAGATGACG	GATAAGCTCA	AAACCCATGT	GAAAGCTCGG	ACAGCTCAAT	360
	TAGCCAAGAT	AAAATGGGTG	ATAAGTGTCG	CTTTCTACGT	ATTGCAGGCT	GCCCTGATGA	420
10	TCTCACTCAT	TTGGAAGTAT	TATTCTGTCC	CTGTGGCTGT	CGTGCCGAGT	AAATGGATAA	480
	CCCTYTAGAC	CGCCTGGTAG	CCTTTCCYAY	TAGAGTAGCA	GGTGGTGTTG	GAATTACTGT	540
15	TGGATTTART	CTGTACAAAT	TGTCCTATTG	TGCTTCACCG	TYCASTGAAC	AGGAGGTGGT	600
15	ACAGCCGGAG	TTAAAAACGG	TTTCCNTTCC	AGTTTAAAAT			640

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(2) INFORMATION FOR SEQ ID NO: 106:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1529 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 106:

GGGCACNAGA TGGAGCTGCC GTAGCGGACC CAGCACAGCC AGGAGCGTCC GGGATGAGCT 60 CAGCCGCGC CGACCACTGG GCGTGGTTGC TGGTGCTCAG CTTCGTGTTT GGATGCAATG 120 TTCTTAGGAT CCTCCTCCCG TCCTTCTCAT CCTTCATGTC CAGGGTGCTG CAGAAGGACG 180 35 CCGACAGGAG TCACAGATGA GAGCGGAGAT CCAGGACATG AAGCAGGAGC TCTCCACAGT 240 300 CAACATGATG GACGAGTTTG CCAGATATGC CAGGCTGGAA AGAAAGATCA ACAAGATGAC 40 GGATAAGCTC AAAACCCATG TGAAAGCTCG GACAGCTCAA TTAGCCAAGA TAAAATGGGT 360 GATAAGTGTC GCTTTCTACG TATTGCAGGC TGCCCTGATG ATCTCACTCA TTTGGAAGTA 420 TTATTCTGTC CCTGTGGCTG TCGTGCCGAG TAAATGGATA ACCCCTCTAG ACCGCCTGGT 45 480 AGCCTTTCCT ACTAGAGTAG CAGGTGGTGT TGGAATTACC TGTTGGATTT TAGTCTGTAA 540 600 CAAAGTTGTC GCTATTGTGC TTCATCCGTT CAGCTGAACA GGAGGATGGA TACAGCCGCG 50 660 AGTAAAAAA CGGATTTCCT CTTCCTAGCT TAAAATCTGA TTTACACTGT TTTGTTTTTT AAGAAACAAA AGTGCATAGT TTAGATTTTT TTTTTGTTGA ATATGTTTGT TCTTGGACTT 720 TATGAGATAG TCTTATAAGA ATCACGATTT TCTACACCTG TCATTGAGCC AAGAAAGTCC 780 55 AGTTTATGAC ACGTATGTAC TAGTGAACAC CGTCCTCGAT CTGTACGAAA TGTGAAATGT 840 TTAGGGACAT CTCCATGCTG TCACTTGTGA TTTGCCCTCT TATGTATTTT GGTCATATTG 900 60

	CCAACTGGAA	AGTCAAAATT	TTCTAACAAC	TTTAAGTAAG	TTCTTTGAAG	ACTTAGTGCT	960
	GTTTTTAATC	CAGTTTAGAA	AGTAACTTAA	TTTTAATACC	RCTACTAAAA	ATTCGAAAAT	1020
5	TTCTTCTTTA	ATCACATTCA	ATATGGTTAA	AAGAACAACA	CTAATTGACA	TTGCGTGGGC	1080
	TTTTTCTCCC	TTTGTTTAAA	ATGTCATTTG	TTGAGCAAGA	GTTGTATAGT	ATTATCTACT	1140
10	TACTTGAGGC	TGTTAATTTT	TCATTACAGT	GTTTTGTAAA	TGTATCCACG	AGACCATGAT	1200
	GCATTGTTTT	GTGCTCAACT	TGTGTTTTGT	ATTTAAAGCA	TTTTGAATGA	AGTGTATTTT	1260
	ATAAGCATTT	AATATTTATG	CTCTTTAGAA	TGGAACACAG	AAAACAAACC	TTATAAGTCC	1320
15	TGATTAATCT	GAACCAATAA	CCTGTGTGGC	CTACAAAGTA	TAATTCTATT	AAATGTTCCT	1380
	TAAAACACTT	TTTTCTAATT	AAAATCTTTG	CAAATGCTTG	TGTAACTTCC	TGCCTTACAG	1440
20	CTACTTGTTT	GCTGTGAGCC	ACCCGCAACT	GACAAGTGGC	TGTTAACTGA	GTCACCATAT	1500
	CCCAGTAAAG	CTGAATTTTC	TCACTAAAA				1529

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(2) INFORMATION FOR SEQ ID NO: 107:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 2435 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 107: 35

55							
	ATGAAGGGTC	GTTGGTGGGA	AAGATGGCGG	CGACTCTGGG	ACCCCTTGGT	CGTGGCAGCA	60
	GTGGCGRCGA	TGTTTGTCGG	CTCGGGATGG	GTCCAGGATG	TTACTCCTTC	TTCTTTTGTT	120
40	GGGGTCTGGG	CAGGGGCCAC	AGCAAGTCGG	GGCGGGTCAA	ACGTTCGAGT	ACTTGAAACG	180
	GGAGCACTCG	CTGTCGAAGC	CCTACCAGGG	TGTGGGCACA	GGCAGTTCCT	CACTGTGGAA	240
45	TCTGATGGGC	AATGCCATGG	TGATGACCCA	GTATATCCGC	CTTACCCCAG	ATATGCAAAG	300
	TAAACAGGGT	GCCTTGTGGA	ACCGGGTGCC	ATGTTTCCTG	AGAGACTGGG	AGTTGCAGGT	360
	GCACTTCAAA	ATCCATGGAC	AAGGAAAGAA	GAATCTGCAT	GGGGATGGCT	TGGCAATCTG	420
50	GTACACAAAG	GRWTCGGATG	CAGCCAGGGC	CTGTNTTTGG	GAAACATGGA	CAAATTTGTG	480
	GGGCTGGGAG	TATTIGTAGA	CACCTACCCC	AATGAGGAGA	AGCAGCAAGA	GCGGGTATTC	540
55	CCCTRCMTCT	CAGCCATGGT	GAACAACGGC	TCCCTCAGCT	ATGATCATGA	GCGGGATGGG	600
	CGGCCTACAG	AGCTGGGAGG	CTGCASAGCC	ATTGTCCGCA	ATCTTCATTA	CGACACCTTC	660
	CTGGTGATTC	GCTACGTCAA	GAGGCATTTR	ACGATAATGA	TGGATATTGA	TGGCAAGCAT	720
60	GAGTGGAGGG	ACTGCATTGA	AGTGCCCGGA	GTCCGCCTGC	CCCGCGGCTA	CTACTTCGGC	780

	ACCTCCTCCA	TCACTGGGGA	TCTCTCAGAT	AATCATGATG	TCATTTCCTT	GAAGTTGTTT	840
5	GAACTGACAG	TGGAGAGAAC	CCCAGAAGAG	GAAAAGCTCC	ATCGAGATGT	GTTCTTGCCC	900
J	TCAGTGGACA	ATATGAAGCT	GCCTGAGATG	ACAGCTCCAC	TGCCGCCCCT	GAGTGGCCTG	960
	GCCCTCTTCC	TCATCGTCTT	TTTCTCCCTG	GGTGTTTTCT	GTATTTGCCA	TAGTCATTGG	1020
10	TATCATACTC	TACAACAAAT	GGCAGGAACA	GAGCCGAAAG	CGCTTCTACT	GAGCCCTCCT	1080
	GCTGCCACCA	CTTTTGTGAC	TGTCACCCAT	GAGGTATGGA	AGGAGCAGGC	ACTGGCCTGA	1140
15	GCATGCAGCC	TGGAGAGTGT	TCTTGTCTCT	AGCAGCTGGT	TGGGGACTAT	ATTCTGTCAC	1200
	TGGAGTTTTG	AATGCAGGGA	CCCCGCATTC	CCATGGTTGT	GCATGGGGAC	ATCTAACTCT	1260
	GGTCTGGGAA	GCCACCCACC	CCAGGGCAAT	GCTGCTGTGA	TGTGCCTTTC	CCTGCAGTCC	1320
20	TTCCATGTGG	GAGCAGAGGT	GTGAAGAGAA	TTTACGTGGT	TGTGATGCCA	AAATCACAGA	1380
	ACAGAATTTC	ATAGCCCAGG	CTGCCGTGTT	GTTTGACTCA	GAAGGCCCTT	CTACTTCAGT	1440
25	TTTGAATCCA	CAAAGAATTA	AAAACTGGTA	ACACCACAGG	CTTTCTGACC	ATCCATTCGT	1500
23	TGGGTTTTGC	ATTTGACCCA	ACCCTCTGCC	TACCTGAGGA	GCTTTCTTTG	GAAACCAGGA	1560
	TGGAAACTTC	TTCCCTGCCT	TACCTTCCTT	TCACTCCATT	CATTGTCCTC	TCTGTGTGCA	1620
30	ACCTGAGCTG	GGAAAGGCAT	TTGGATGCCT	CTCTGTTGGG	GCCTGGGGCT	GCAGAACACA	1680
	CCTGCGTTTC	ACTGGCCTTC	ATTAGGTGGC	CCTAGGGAGA	TGGCTTTCTG	CTTTGGATCA	1740
35	CTGTTCCCTA	GCATGGGTCT	TGGGTCTATT	GGCATGTCCA	TGGCCTTCCC	AATCAAGTCT	1800
55	CTTCAGGCCC	TCAGTGAAGT	TTGGCTAAAG	GTTGGTGTAA	AAATCAAGAG	AAGCCTGGAA	1860
	GACATCATGG	ATGCCATGGA	TTAGCTGTGC	AACTGACCAG	CTCCAGGTTT	GATCAAACCA	1920
40	AAAGCAACAT	TTGTCATGTG	GTCTGACCAT	GTGGAGATGT	TTCTGGACTT	GCTAGAGCCT	1980
	GCTTAGCTGC	ATGTTTTGTA	GTTACGATTT	TTGGAATCCC	ACTTTGAGTG	CTGAAAGTGT	2040
45	AAGGAAGCTT	TCTTCTTACA	CCTTGGGCTT	GGATATTGCC	CAGAGAAGAA	ATTIGGCTTT	2100
73	TTTTTTNCTT	AATGGACAAG	AGACAGTTGC	TGTTCTCATG	TTCCAAGTCT	GAGAGCAACA	2160
	GACCCTCATC	ATCTGTGCCT	GGAAGAGTTC	ACTGTCATTG	AGCAGCACAG	CCTGAGTGCT	2220
50	GGCCTCTGTC	AACCCTTATT	CCACTGCCTT	T ATTTGACAAG	GGGTTACATO	CTGCTCACCT	2280
	TACTGCCCTC	GGATTAAATO	AGTTACAGGG	CAGAGTCTCC	TTGGAGGGC	TGGAACTCTG	2340
55	AGTCCTCCT	A TGAACCTCTC	TAGCCTAAA	GAAATTCTTA	A AAATCACCGA	A TGGAACCAAA	2400
33	AAAAAAAA	AAAAAAAA	AAAAAAAA	A AAAAN			2435

	(2) INFORMATION FOR SEQ ID NO: 108:						
5	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 805 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear						
10	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 108:						
10	ATGAAACTTA AGAATTGAAT TGGAAAGACT TCTCAAAGAG AATTGTATGT AACGATGTTG	60					
	TATTGATTTT TAAGAAAGTA ATTTAATTTG TAAAACTTCT GCTCGTTTAC ACTGCACATT	120					
15	GAATACAGGT AACTAATTGG AAGGAGAGGG GAGGTCACTC TTTTGATGGT GGCCCTGAAC	180					
	CTCATTCTGG TTCCCTGCTG CGCTGCTTGG TGTGACCCAC GGAGGATCCA CTCCCAGGAT	240					
20	GACGTGCTCC GTAGCTCTGC TGCTGATACT GGGTCTGCGA TGCAGCGGCG TGAGGCCTGG	300					
	GCTGGTTGGA GAAGGTCACA ACCCTTCTCT GTTGGTCTGC CTTCTGCTGA AAGACTCGAG	360					
	AACCAACCAG GGAAGCTGTC CTGGAGGTCC CTGGTCGGAG AGGGACATAG AATCTGTGAC	420					
25	CTCTGACAAC TGTGAAGCCA CCCTGGGCTA CAGAAACCAC AGTCTTCCCA GCAATTATTA	480					
	CAATTCTTGA ATTCCTTGGG GATTTTTTAC TGCCCTTTCA AAGCACTTAA GTGTTAGATC	540					
30	TAACGTGTTC CAGTGTCTGT CTGAGGTGAC TTAAAAAATC AGAACAAAAC TTCTATTATC	600					
50	CAGAGTCATG GGAGAGTACA CCCTTTCCAG GAATAATGTT TTGGGAAACA CTGAAATGAA	660					
	ATCTTCCCAG TATTATAAAT TGTGTATTTA AAAAAAAGAA ACTTTTCTGA ATGCCTACTG	720					
35	GCGGTGTATA CCAGGCAGTG TGCCAGTTTA AAAAGATGAA AAAGAATAAA AACTTTTGAG	780					
	GAACAAAAA AAAAAAAAA AAATT	805					
40							
	(2) INFORMATION FOR SEO ID NO: 109:						
45	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1166 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear						
50	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 109:						
	GGCACGAGAG GCGCCAGTCG CAGGTGTGCT GCTGAGGCGT GAGAATGGCG TCCCGCGGCC	60					
c c	GGCGTCCGGA GCATGGCGGA CCCCCAGAGC TGTTTTATGA CGAGACAGAA GCCCGGAAAT	120					
55	ACGTTCGCAA CTCACGGATG ATTGATATCC AGACCAGGAT GGCTGGGCGA GCATTGGAGC	180					
	TTCTTTATCT GCCAGAGAAT AAGCCCTGTT ACCTGCTGGA TATTGGCTGT GGCACTGGGC	240					
60	TGAGTGGAAG TTATCTGTCA GATGAAGGGC ACTATTGGGT GGGCCTGGAT ATCAGCCCTG	300					

	CCATGCTGGA	TGAGGCTGTG	GACCGAGAGA	TAGAGGGAGA	CCTGCTGCTG	GGGGATATGG	360
5	GCCAGGGCAT	CCCATTCAAG	CCAGGCACAT	TTGATGGTTG	CATCAGCATT	TCTGCTGTGC	420
3	AGTGGCTCTG	TAATGCTAAC	AAGAAGTCTG	AAAACCCTGC	CAAGCGCCTG	TACTGCTTTT	480
	TTGCTTCTCT	TTTTTCTGTT	CTCGTCCGGG	GATCCCGAGC	TGTCCTGCAG	CTGTACCCTG	540
10	AGAACTCAGA	GCAGTTGGAG	CTGATCACAA	CCCAGGCCAC	AAAGGCAGGC	TTCTCCGGTG	600
	GCATGGTGGT	AGACTACCCT	AACAGTGCCA	AAGCAAAGAA	ATTCTACCTC	TGCTTGTTTT	660
15	CTGGGCCTTC	GACCTTTATA	CCAGAGGGGC	TGAGTGAAAA	TCAGGATGAA	GTTGAACCCA	720
15	GGGAGTCTGT	GTTCACCAAT	GAGAGGTTCC	CATTAAGGAT	GTCGAGGCGG	GGAATGGTGA	780
	GGAAGAGTCG	GGCATGGGTG	CTGGAGAAGA	AGGAGCGGCA	CAGGCGCCAG	GGCAGGGAAG	840
20	TCAGACCTGA	CACCCAGTAC	ACCGGCCGCA	AGCGCAAGCC	CCGCTTCTAA	GTCACCACGC	900
	GGTTCTGGAA	AGGCACTTGC	CTCTGCACTT	TTCTATATTG	TTCAGCTGAC	AAAGTAGTAT	960
25	TTTAGAAAAG	TTCTAAAGTT	ATAAAAATGT	TTTCTGCAGT	AAAAAAAAG	TTCTCTGGGC	1020
23	CGGGCGTGGT	GGCTCACANC	TGTAATCCCA	GCACCTTGGG	AGGCTGAGGT	GGGAGGATCA	1080
	TTTGAGGCCA	GGAGTTTGAG	ACCTGCCTGG	GCAACATAAT	GAAACTTCCT	TTCCAGGGAG	1140
30	AAAAAAAAA	AAAAAAAAA	ACTCGA				1166

35 (2) INFORMATION FOR SEQ ID NO: 110:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 586 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 110:

45	AGAGCGGACG	AAGCTGGATA	ACAGGGGACC	GATGATGTGG	CGACCATCAG	TTCTGCTGCT	60
	TCTGTTGCTA	CTGAGGCACG	GGGCCCAGGG	GAAGCCATCC	CCAGACGCAG	GCCCTCATGG	120
50	CCAGGGGAGG	GTGCACCAGG	CGGCCCCCT	GAGCGACGCT	CCCCATGATG	ACGCCCACGG	180
30	GAACTTCCAG	TACGACCATG	AGGCTTTCCT	GGGACGGGAA	GTGGCCAAGG	AATTCGACCA	240
	ACTCACCCCA	GAGGAAAGCC	AGGCCCGTCT	GGGGCGGATC	GTGGACCGCA	TGGACCGCGC	300
55	GGGGACGGC	GACGGCTGGG	TGTCGCTGGC	CGAGCTTCGC	GCGTGGATCG	CGCACACGCA	360
	GCAGCGGCAC	ATACGGGACT	CGGTGAGCGC	GGCCTGGGAC	ACGTACGACA	CGGACCGCGA	420
60	CGGGCGTGTG	GGTTGGGAGG	AGCTGCGCAA	CGYCACCTAT	GGCCACTASG	SGCCCGKTGA	480
00							

260

	AGAATITCAT	GACGTGGAGG	ATGCAGAGAC	YTACAAAAAG	ATGCTGGYTC	GGGACGAGCG	540		
	GCGTTTCCGG	GTGGCCGACC	AGGATGGGGA	CTCGATGGCC	ACTCGA		586		
5									
	(2) INFORMA	ATION FOR SE	Q ID NO: 11	L1:					
10	(i)	(B) TYP (C) STR	HARACTERIST: GTH: 1134 b E: nucleic ANDEDNESS: DLOGY: line	ase pairs acid double					
15	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 111:								
	ACCCATTGAG	CAGAAGGAGG	CCAGGTGGGA	AAGCTCCTGG	GAAGAGCAGC	CAGACTGGAC	60		
20	ACTGGGCTGC	TTGAGTCCTG	AGTCACAATT	CAGAATTCCT	GGGCTCCCTG	GGTGCATTCT	120		
	ATCATTCCAG	TTGAAAGTTT	GCTTCCTTCC	AGTCATGTGG	CTCTTCATTC	TACTCTCCTT	180		
25	GGCTCTCATT	TCAGATGCCA	TGGTCATGGA	TGAAAAGGTC	AAGAGAAGTT	TGTGCTGGAC	240		
20	ACGGCTTCTG	CCATCTGCAA	CTACAATGCC	CAYTACAAGA	ATCACCCCAA	ATACTGGTGC	300		
	CGAGGYTATT	TCCGTGAYTA	CTGCAACATC	ATCGCCTTCT	CCCCTAACAG	CACCAATCAT	360		
30	GTGGCCCTGA	AGGACACAGG	GAACCAGCTC	ATTGTCACTA	TGTCCTGCCT	GAACAAANAA	420		
	GACACGGGCT	GGTACTGGTG	TGGCATÇCAR	CGGGACTTTG	CMAGGGATGA	CATGGATTTT	480		
35	ACAGAGCTGA	TTGTAACTGA	CGACAAAGGA	ACCCTGGCCA	ATGACTTTTG	GTCTGGGAAA	540		
	GACCTATCAG	GCAACAAAAC	CAGAAGCTGC	AAGGCTCCCA	AAGTTGTCCG	CAAGCTGACC	600		
	GCTCCAGGAC	GTCCATTCTC	ATCATTTGCA	TACTGATCAC	GGGTTTGGGA	ATCATCTCTG	660		
40	TAATCAGTCA	TTTGACCAAA	AGGAGGAGAA	GTCAAAGGAA	TAGAAGGGTA	GGCAACACTT	720		
	TGAAGCCCTT	CTCGCGTGTC	CTGACTCCAA	AGGAAATGGC	TCCTACTGAA	CAGATGTGAC	780		
45	TGAAGWITTT	TTTAATTTAG	TTNCATAAAG	TGATGNCTAC	AACAGAWTAA	TCACCCATGA	840		
	CAACTGGCCC	CACACCTCAG	AGACTGATTC	TGATCTCCCA	GGAATTCTGA	AGGACCCTCT	900		
	ATCCTTGACA	ACAATCATTT	GCAGCCAGGT	AGCAACGGCR	GTAGTCAGAG	GAGCTATGAT	960		
50	AGACCACACC	CAAGCAAGGC	TGCCCTCAAA	TAACATCTCA	AGATCTTAGT	TCTTATGCAT	1020		
	TCCATCAGTC	AGAAGTGAAG	AAGAGGTGGA	GAATCTKGAT	TGGGGACCAG	GAAATCACTT	1080		
55	GTATTTIGTT	AGCCAATAAA	TI . JTAGCCA	GTGTTGAATG	АААААААА	AAAA	1134		

(2) INFORMATION FOR SEQ ID NO: 112:

(A) LENGTH: 1333 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double
(D) TOPOLOGY: linear

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 112:

10	CACTTTAAAG CTCTGCTGAG GGAGTTCGGA GCCCAGGCTT TCAGGCGACC TCTGCCCTCC	60
10	CTGCCTCTCC TCACCCTCCC TCTCTTCCTG CAGGGCCTGG GAAGGGCTTT GAGGGAGCCT	120
	GGGAGCCATG TGAAGAGGGG CACGCCTGGG CTGTCCCACA GTTTAGATCC AGTTGGAGGT	180
15	TCTCCCTGGC TCCTGCAGGC CTGCGGGGAT CTCTCCCCAC TTCAGGCCTC CGGCAGCTGC	240
	CTGCCCTCTT GTCTGTGCTT CAGCCCTGCA CAAAAGCAGC TTGGTGACAC CACTCAGCCA	300
•	CCCAGAGTAC GTGTTTACAG GCTTTCCAGA TCACCTTCCT GTGGGGTGAA CGTAATGAGG	360
20	CGGGGCTGGT CCTTGGAATT TCCCCTGGAA AATGGTAACA GACTCCATCC TTGACCCGGG	420
	GATGAGCATG AAGGCATTGT CCCAAAGGCA GAGGCCACCG TGGTAGGAAT TCCACCAAGG	480
25	CCAGAAGGGA AAAAGGAAGA ACCCACCGTG TCTGGCTGTG CGGGCCCTGG GGAGGGTCGT	540
	GAGTGCAGCC CCTCTCTACT TCYGTGCCTT TGTAAAACGT GTAGATAACC GCAGTGGTTG	600
20	GCTGAGCCAA GAACTCTCCT AAATCAGTGG CTTTCTCCCC ACCCCTTGCT GGGGAGTCAT	660
30	TTTTAAAAAA ATCTGTGGGA TATAAAATTG GCCTCCTGCT GCTTCAGCCT ACCTCTCCCT	720
	CTGCTGACTT AATGTCGTGA TTCTGTTTCT TCAGATATTT AAGGCTGTTA GGTTGTGTGA	780
35	GCCTTGAAGT GTGTGTGTGT GTCCCAGCGA CTGTCCACTG TCCAGGAGAT GCATGTCTTT	840
	GTATTGGAGA TATTTCTGTA ACTCATTCTC TTGGTGCTCA CGATTGCCAT GGCCATAGGG	900
40	CCACAGTGCC GTATCTGCTG CAGACATGAT TGTTTCTTGT TCTAGAGGTT TTCTTGTTTT	960
40	CGAATCTTGC CTGATGAATC CAGCCAGACC AAGGGGCCTA GATTTGACCT CTGTCCTGGG	1020
	CTCCTGGGCC AGGTGCAGGA ACATCTGAGG CCACTCTGCT GGCCACCTCC AGTGGGTGCT	1080
45	GACCACAGGA TGGGCTTTGT TTACACTCAT TTTCACCCTG ATTCTTGCCC CCACTTTCAT	1140
	AAAAGAAACT TCAAAATGCT GACGCTTTGG AGAGTAAGAA AATCAATCTT GGCTGGGCAC	1200
50	GGTGGCTCCT GCCTGTGATC CTAGCACTTT GGGAGGCTGA AGCTGAAGGA TCACTTGAGC	1260
50	TCAGGAGTTG GAGACCAACC CTGGCAACAT AACAAGACCC TGTCTCTACA AAAAAAAAAA	1320
	AAAAAAACT CGA	1333

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(2) INFORMATION FOR SEQ ID NO: 113:

60 (i) SEQUENCE CHARACTERISTICS:

WO 98/42738

5	(A) LENGTH: 1015 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	۰
3	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 113:	
	GGCACGAGCG GCACGAGCGG CACGAGGTGA CTTCAAGTGT CGGATCTTTT CAGCCTACAT	60
10	CAAGGAGGTG GAGGAACGGC CGGCACCCAC CCCGTGGGCT CCAAGATGCC CTTTGGGGAA	120
	CTGATGTTCG AATCCAGCAG TAGCTGCGGC TGGGTACATG GCGTCTGTTT CTCAGCCAGC	180
15	GGGAGCCGCG TGGCCTGGGT AAGCCACGAC AGCACCGTCT GCCTGGCTGA TGCCGACAAG	240
13	AAGATGGCCG TCGCGACTCT GGCCTCTGAA ACACTACCAC TGCTGGCGCT GACCTTCATC	300
	ACAGACAACA GCCTGGTGGC AGCGGGCCAC GACTGCTTCC CGGTGCTGTT CACCTATGAC	360
20	GCCGCCGCGG GGATGCTGAG CTTCGGCGGG CGGCTGGACG TTCCTAAGCA GAGCTCGCAG	420
	CGTGGCTTGA CGGCCCGCGA GCGCTTCCAG AACCTGGACA AGAAGGCGAG CTCCGAGGGT	480
25	GGCACGGCTG CGGGCGCGGG CCTAGACTCG CTGCACAAGA ACAGCGTCAG CCAGATCTCG	540
	GTGCTCAGCG GCGGCAAGGC CAAGTGCTCG CAGTTCTGCA CCACTGGCAT GGATGGCGGC	600
	ATGAGTATCT GGGATGTGAA GAGCTTGGAG TCAGCCTTGA AGGACCTCAA GATCAAATGA	660
30	CCTGTGAGGA ATATGTTGCC TTCATCCTAG CTGCTGGGGA AGCGGGGAGA GGGGTCAGGG	720
	AGGCTAATGG TTGCTTTGCT GAATGTTTCT GGGGTACCAA TACGAGTTCC CATAGGGGCT	780
35	GCTCCCTCAA AAAGGGAGGG GACAGATGGG GAGCTTTTCT TACCTATTCA AGGAATACGT	840
	GCCTTTTTCT TAAATGCTTT CATTTATTGA AAAAAAAAA AAATGCCCCC AAAGCACTAT	900
	GCTGGTCATG AACTGCTTCA AAATGTGGAG GTAATAAAAT GCAACTGTGT AAAAAAAAA	960
40	AAAAAAAAA AAAAAAAAA AAAAAAAAA AAAAAAAA	1015
45	(2) INCOMMENDATION FOR SUC. ID NO. 114	
43	(2) INFORMATION FOR SEQ ID NO: 114:	
50	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1076 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 114:	
55	GGCACGAGGG GAAAGCCATG CTCCCAGGAC TCCTTCCTTG CAGCCTTAAA TCGGTCTGTA	60
	CGGAAAATTC CGCGCCTTAG AAACCCACGC TTGGGTGTAA CTTATTATTG TTCTTCCTGA	120
60	CCTACTTCCT GTTTATCACT TCCGGGTTCA TCATTTTGGC ATTTCGGTGA TCGGGTTGGA	180

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	ACTATTGAAG CCCGCTTTCA GGTTCTTTTC CCCATTTTCC CTTTGAAAGG AAGACTTCTG	240
	GCTTCTCCTA AATCTCCGTT CTCTGGGTAA GGGGAGTCCA AGCCTCTGTC ATGAGGAACG	300
5	GAAATGCGAG GGCCTCGGGT GTTACTCTAA AATCCGCCCT CAGCTTGCAC GCCGGAAGCT	360
	GCGATTCCTG CAGCGGAAGA GGCGTGATCT GGCCTTCGAC TCGCTATGTC CACTAACAAT	420
	ATGTCGGACC CACGGAGGCC GAACAAAGTG CTGAGGTACA AGCCCCCGCC GAGCGAATGT	480
10	AACCCGGCCT TGGACGACCC GACGCCGGAC TACATGAACC TGCTGGGCAT GATCTTCAGC	540
	ATGTGCGGCC TCATGCTTAA GCTGAAGTGG TGTGCTTGGG TCGCTGTCTA CTGCTCCTTC	600
15	ATCAGCTTTG CCAACTCTCG GAGCTCGGAG GACACGAAGC AAATGATGAG TAGCTTCATG	660
	CTGTCCATCT CTGCCGTGGT GATGTCCTAT CTGCAGAATC CTCAGCCCAT GACGCCCCCA	720
20	TGGTGATACC AGCCTAGAAG GGTCACATTT TGGACCCTGT CTATCCACTA GGCCTGGGCT	780
20	TTGGCTGCTA AACCTGCTGC CTTCAGCTGC CATCCTGGAC TTCCCTGAAT GAGGCCGTCT	840
	CGGTGCCCC AGCTGGATAG AGGGAACCTG GCCCTTTCCT AGGGAACACC CTAGGCTTAC	900
25	CCCTCCTGCC TCCCTTCCCC TGCCTGCTGC TGGGGGAGAT GCTGTCCATG TTTCTAGGGG	960
	TATTCATTTG CTTTCTCGTT GAAACCTGTT GTTAATAAAG TTTTTCACTC TGAAAAAAAA	1020
30	AAAAAAAAAA RAAAACNCGN GGGGGGCCC GGAACCCAAT TCSCCGGATA GTGAGT	1076
35 40	(2) INFORMATION FOR SEQ ID NO: 115: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1487 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 115:	
45	CCGCTGCTGA TAACTATGGC ATCCCCCGGG CCTGCAGGAA TTCGGCACGG AGCTACGGCG	60
13	CCGCCTGGCT CCTGCTGNCA CCTGCAGGCT CGTCGCGGGT GGAGCCCACC CAAGACATCA	120
	GCATCAGCGA CCAGCTGGGG GGCCAGGACG TGCCCGTGTT CCGGAACCTG TCCCTGCTGG	180
50	TGGTGGGTGT CGGCGCCGTG TTCTCACTGC TATTCCACCT GGGCACCCGG GAGAGGCGCC	240
	GGCCGCATGC GGASGAGCCA GGCGAGCACA CCCCCCTGTT GGCCCCTGCC ACGGCCCAGC	300
55	CCCTGCTGCT CTGGAAGCAC TGGCTCCGGG AGCSGGCTTT CTACCAGGTG GGCATACTGT	360
55	ACATGACCAC CAGGCTCATC GTGAACCTGT CCCAGACCTA CATGGCCATG TACCTCACCT	420
	ACTCGCTCCA CCTGCCCAAG AAGTTCATCG CGACCATTCC CCTGGTGATG TACCTCAGCG	480

60 gettettgte eteetteete atgaageeea teaacaagtg cattgggagg aacatgaeet

	ACTTCTCAGG	CCTCCTGGTG	ATCCTGGCCT	TTGCCGCCTG	GGTGGCGCTG	GCGGAGGGAC	600
5	TGGGTGTGGC	CGTGTACGCA	GCGGCTGTGC	TGCTGGGTGC	TGGCTGTGCC	ACCATCCTCG	660
5	TCACCTCGCT	GGCCATGACG	GCCGACCTCA	TCGGTCCCCA	CACGAACAGC	GGAGCKTTCG	720
	TGTACGGCTC	CATGAGCTTC	TTGGATAAGG	TGGCCAATGG	GCTGGCAGTC	ATGGCCATCC	780
10	AGAGCCTGCA	CCCTTGCCCC	TCAGAGCTCT	GCTGCAGGGC	CTGCGTGAGC	TTTTACCACT	840
	GGGCGATGGT	GGCTGTGACG	GCCGCCTGG	GCGTGGCCGC	TGCCCTGTGT	CTCTGTAGCC	900
15	TCCTGCTGTG	GCCGACCCGC	CTGCGACGCT	GATGAGACCT	GCACGCANTG	GCTCACAGCA	960
13	GCACGATTTG	TGACAGCCCG	AGGCGGAGAA	CACCGAACAC	CCAGTGAAGG	TGAGGGGATC	1020
	AGCACGGCGC	GGCCACCCAC	GCACCCACGC	GCTGGAATGA	GACTCAGCCA	CAAGGAGGTG	1080
20	CGAAGCTCTG	ACCCAGGCCA	CAGTGCGGAT	GCACCTTGAG	GATGTCACGC	TCAGTGAGAG	1140
	ACACCAGACA	CAGAAGGGTA	CGCTGTGATC	CCACTTCTAT	GAAATGTCCA	GGACAGACCA	1200
25	ATCCACAGAA	TCAGGGAGAG	GATTCGTGGG	TGCCGGGACT	GGGGAGGGG	ACCTGGGGGT	1260
	GACTAGGTGA	CATAATGGGG	ACAGGGCTGC	CTTCTGGGTG	ATGAGAATGT	TCTGGAATCA	1320
	GATGGGATGG	CTGCACGGCG	TGGTGAAGGT	ACTGAACGCC	ACCTCACTGT	AAGACGGTAG	1380
30	ATTTTGTATT	TTACCACAAT	AAACAAAACA	AAACAAAACC	AAAAAAAAA	AAAAAAAA	1440
	AAAAAAAAGG	AATTCGATAT	CAAGCTTATC	GATACCGTCG	ACCTCGA		1487
35							
	(2) INFORM	ATION FOR S	EQ ID NO: 1	16:			
40	(i)		HARACTERIST GTH: 1350 b	ase pairs			
		,	ANDEDNESS: OLOGY: line				
45	(xi) SEQUENCE	DESCRIPTION	: SEQ ID NO	: 116:		
	GGCACGAGTG	CGCANGCGTG	GGGCTCTCTC	CTTGTCAGTC	GGCGCCGCGT	GCGGGCTGGT	60
••	GGCTCTGTGG	CAGCGGCGGC	GGCAGGACTC	CGGCACTATG	AGCGGCTTCA	GCACCGAGGA	120
50	GCGCGCCGCG	CCTTCTCCCT	GGAGTACCGA	GTCTTCCTCA	AAAATGAGAA	AGGACAATAT	180
	ATATCTCCAT	TTCATGATAT	TCCAATTTAT	GCAGATAAGG	ATGTGTTTCA	CATGGTAGTT	240
55	GAAGTACCAC	GCTGGTCTAA	TGCAAAAATG	GAGATTGCTA	CAAAGGACCC	TTTAAACCCT	300
	ATTAAACAAG	ATGTGAAAAA	AGGAAAACTT	CGCTATGTTG	CGAATTTGTT	CCCGTATAAA	360
60	GGATATATCT	GGAACTATGG	TGCCATCCCT	CAGACTTGGG	AAGACCCAGG	GCACAATGAT	420
60							

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	AAACATACTG GCTGTTGTGG TGACAATGAC CCAATTGATG TGTGTGAAAT TGGAAGCAAG	480
	GTATGTGCAA GAGGTGAAAT AATTGGCGTG AAAGTTCTAG GCATATTGGC TATGATTGAC	540
5	GAAGGGGAAA CCGACTGGAA AGTCATTGCC ATTAATGTGG ATGATCCTGA TGCAGCCAAT	600
	TATAATGATA TCAATGATGT CAAACGGCTG AAACCTGGCT ACTTAGAAGC TACTGTGGAC	660
10	TGGTTTAGAA GGTATAAGGT TCCTGATGGA AAACCAGAAA ATGAGTTTGC GTTTAATGCA	720
10	GAATTTAAAG ATAAGGACTT TGCCATTGAT ATTATTAAAA GCACTCATGA CCATTGGAAA	780
	GCATTAGTGA CTAAGAAAAC GAATGGAAAA GGAATCAGTT GCATGAATAC AACTTTGTCT	840
15	GAGAGCCCCT TCAAGTGTGA TCCTGATGCT GCCAGAGCCA TTGTGGATGC TTTACCACCA	900
	CCCTGTGAAT CTGCCTGCAC AGTACCAACA GACGTGGATA AGTGGTTCCA TCACCAGAAA	960
20	AACTAATGAG ATTTCTCTGG AATACAAGCT GATATTGCTA CATCGTGTTC ATCTGGATGT	1020
20	ATTAGAAGTA AAAGTAGTAG CTTTTCAAAG CTTTAAATTT GTAGAACTCA TCTAACTAAA	1080
	GTAAATTCTG CTGTGACTAA TCCAATATAC TCAGAATGTT ATCCATCTAA AGCATTTTTC	1140
25	ATATCTCAAC TAAGATAACT TTTAGCACAT GCTTAAATAT CAAAGCAGTT GTCATTTGGA	1200
	AGTCACTTGT GAATAGATGT GCAAGGGGAG CACATATTGG ATGTATATGT TACCATATGT	1260
20	TAGGAAATAA AATTATTTTG CTGAAAAAAA AAAAAAAAAA	1320
30	CCCCATTIGG CCCTITGGGG GGNGGTTTTA	1350
35	(2) INFORMATION FOR SEQ ID NO: 117:	
40	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 2527 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear 	
45	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 117:	
43	CTCTTGCTAC CTTCCCGGCG CAGAGAACCC CGGCTGCTCA GCGCGCTCCG GGGTCATGGA	60
	GATCCCCGGG AGCCTGTGCA AGAAAGTCAA GCTGAGCAAT AACGCGCAGA ACTGGGGAAT	120
50	GCAGAGAGCA ACCAATGTCA CCTACCAAGC CCATCATGTC AGCAGGAACA AGAGAGGTCA	180
	GGTGGTGGG ACCAGAGGTG GCTTTCGTGG TTGCACAGTT TGGCTAACAG GCTTGTCTGG	240
<i>e e</i>	AGCGGGAAAG ACTACTGTGA GCATGGCCTT GGAGGAGTAC CTGGTTTGTC ATGGTATTCC	300
55	ATGCTACACT CTGGATGGTG ACAATATTCG TCAAGGTCTC AATAAAAATC TTGGCTTTAG	360
	TCCTGAAGAC AGAGAAGAGA ATGTTCGACG CATCGCAGAA GTTGCTAAAC TGTTTGCAGA	420

60 TGCTGGCTTA GTGTGCATCA CAAGTTTCAT ATCACCTTAC ACTCAGGATC GCAACAATGC

	AAGGCAAATT	CATGAAGGTG	CAAGTTTACC	GTTTTTTGAA	GTATTTGTTG	ATGCTCCTCT	540
5	GCATGTTTGT	GAACAGAGGG	ATGTCAAAGG	ACTCTACAAA	AAAGCCCGGG	CAGGAGAAAT	600
3	TAAAGGTTTC	ACTGGGATCG	ATTCTGAATA	TGAAAAGCCA	GAGGCCCCTG	AGTIGGTGCT	660
	GAAAACAGAC	TCCTGTGATG	TAAATGACTG	TGTCCAGCAA	GTTGTGGAAC	TTCTACAGGA	720
10	ACGGGATATT	GTACCTGTGG	ATGCATCTTA	TGAAGTAAAA	GAACTATATG	TGCCAGAAAA	780
	TAAACTTCAT	TTGGCAAAAA	CAGATGCGGA	AACATTACCA	GCACTGAAAA	TTAATAAAGT	840
15	GGATATGCAG	TGGGTGCAGG	TTTTGGCAGA	AGGTTGGGCA	ACCCCATTGA	ATGGCTTTAT	900
	GAGAGAGAGG	GAGTACTTGC	AGTGCCTTCA	TTTTGATTGT	CTTCTGGATG	GAGGTGTCAT	960
	TAACTTGTCA	GTACCTATAG	TTCTGACTGC	GACTCATGAA	GATAAAGAGA	GGCTGGACGG	1020
20	CTGTACAGCA	TTTGCTCTGA	TGTATGAGGG	CCGCCGTGTG	GCCATTCTTC	GCAATCCAGA	1080
	GTTTTTTGAG	CACAGGAAAG	AGGAGCGCTG	TGCCAGACAG	TGGGGAACGA	CATGCAAGAA	1140
25	CCACCCCTAT	ATTAAGATGG	TGATGGAACA	AGGAGATTGG	CTGATTGGAG	GAGATCTTCA	1200
	AGTCTTGGAT	CGAGTTTATT	GGAATGATGG	TCTTGATCAG	TATCGTCTTA	CTCCTACTGA	1260
	GCTAAAGCAG	AAATTTAAAG	ATATGAATGC	TGATGCTGTC	TTTGCATTTC	AACTACGCAA	1320
30	CCCAGTGCAC	AATGGACATG	CCCTGTTAAT	GCAGGATACC	CATAAGCAAC	TTCTAGAGAG	1380
	GGGCTACCGG	CGCCCTGTCC	TCCTCCTCCA	CCCTCTGGGT	GGCTGGACAA	AGGATGACGA	1440
35	TGTTCCTTTG	ATGTGGCGTA	TGAAGCAGCA	TGCTGCAGTG	TTGGAGGAAG	GAGTTCTGAA	1500
	TCCTGAGACG	ACAGTGGTGG	CCATCTTCCC	ATCTCCCATG	ATGTATGCTG	GACCAACTGA	1560
	GGTCCAGTGG	CATTGCAGAG	CACGGATGGT	TGCAGGAGCC	AACTTTTACA	TTGTTGGACG	1620
40	AGACCCTGCT	GGCATGCCTC	ATCCAGAAAC	AGGGAAGGAT	CTTTATGAGC	CAAGTCATGG	1680
	TGCCAAAGTG	CTGACGATGG	CCCCTGGTTT	AATCACTTTG	GAAATAGTTC	CCTTTCGAGT	1740
45	TGCAGCTTAC	AACAAGAAAA	AGAAGCGTAT	GGACTACTAT	GACTCTGAAC	ACCATGAAGA	1800
	CTTTGAATTT	ATTTCAGGAA	CACGAATGCG	CAAACTTGCT	CGAGAAGGCC	AGAAACCACC	1860
	TGAAGGTTTC	ATGGCTCCCA	AGGCTTGGAC	CGTGCTGACA	GAATACTACA	AATCCTTGGA	1920
50	GAAAGCTTAG	GCTGTTAACC	CAGTCACTCC	ACCTTTGACA	CATTACTAGT	AACAAGAGGG	1980
	GACCACATAG	TCTCTGTTGG	CATTTCTTTG	TGGTGTCTGT	CTGGACATGC	TTCCTAAAAA	2040
55	CAGACCATTT	TCCTTAACTT	GCATCAGTTT	TGGTCTGCCT	TATGAGTTCT	GTTTTGAACA	2100
	AGTGTAACAC	ACTGATGGTT	TTAATGTATC	TTTTCCACTT	ATTATAGTTA	TATTCCTACA	2160
	ATACAATTTT	AAAATTGTCT	TTTTATATTA	TATTTATGCT	TCTGTGTCAT	GATTTTTTCA	2220
60	AGCTGTTATA	TTAGTTGTAA	CCAGTAGTAT	ТСАСАТТААА	TCTTGCTTTT	TTTCCCCTTA	2280

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	AAAAAAGAAA AAAATTACCA AACAATAAAC TTGGCTAGAC CTTGTTTTGA GGATTTTACA	2340
_	AGACCTTTGT AGCGATTAGA TTTTTTTCT ACATTGAAAA TAGAAACTGC TTCCTTTCTT	2400
5	CTTTCCAGTC AGCTATTGGT CTTTCCAGCT GTTATAATCT AAAGTATTCT TATGATCTGT	2460
	GTAAGCTCTG AATGAACTTC TTTACTCAAT AAAATTAATT TTTTGGCTTC TTAAAAAAAA	2520
10	ааааааа	2527
15	(2) INFORMATION FOR SEQ ID NO: 118:	
15		
	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 1098 base pairs	
20	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
20	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 118:	
25	CGCATCACAG ACAACCCAGA AGGAAAATGG TTGGGCAGAA CAGCAAGGGG TTCATATGGC	60
	TATATTAAAA CAACTGCTGT AGAGATTNNC TATGATTCTT TGAAACTGAA AAAAGACTCT	120
•	CTTGGTGCCC CTTCAAGACC TATTGAAGAT GACCAAGAAG TATATGATGA TGTTGCAGAG	180
30	CAGGATGATA TTAGCAGCCA CAGTCAGAGT GGAAGTGGAG GGATATTCCC TCCACCACCA	240
	GATGATGACA TTTATGATGG GATTGAAGAG GAAGATGCTG ATGATGGTTT CCCTGCTCCT	300
35	CCTAAACAAT TGGACATGGG AGATGAAGTT TACGATGATG TGGATACCTC TGATTTCCCT	360
	GTTTCATCAG CAGAGATGAG TCAAGGAACT AATGTTGGAA AAGCTAAGAC AGAAGAAAAG	420
40	GACCTTAAGA AGCTAAAAAA GCAGRAAAAA GAARAAAAAG ACTTCAGGAA AAAATTTAAA	480
40	TATGATGGTG AAATTAGAGT CCTATATTCA ACTAAAGTTA CAACTTCCAT AACTTCTAAA	540
	AAGTGGGGAA CCAGAGATCT ACAGGTAAAA CCTGGTGAAT CTCTAGAAGT TATACAAACC	600
45	ACAGATGACA CAAAAGTTCT CTGCAGAAAT GAAGAAGGGA AATATGGTTA TGTCCTTCGG	660
	AGTTACCTAG CGGACAATGA TGGAGAGATC TATGATGATA TTGCTGATGG CTGCATCTAT	720
50	GACAATGACT AGCACTCAAC TTTGGTCATT CTGCTGTGTT CATTAGGTGC CAATGTGAAG	780
50	TCTGGATTTT AATTGGCATG TTATTGGGTA TCMAGAAAAT TAATGCACAR AACCACTTAT	840
	TATCATTTGT TATGAAATCC CAATTATCTT TACAAAGTGT TTAAAGTTTG AACATAGAAA	900
55	ATAATCTCTC TGCTTAATTG TTATCTCAGA AGACTACATT AGTGAGATGT AAGAATTATT	960
	AAATATTCCA TTTCCGCTTT GGCTACAATT ATGAAGAAGT TGAAGGTACT TCTTTTAGAC	1020

GGGGGCCCGG TACCCAAT 1098

5

10

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(2) INFORMATION FOR SEQ ID NO: 119:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 1679 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDNESS: double

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 119:

TCGACCCACG CGTCCGGCGA GATCCCTACC GCAGTAGCCG CCTCTGCCGC CGCGGAGCTT 60 CCCGAACCTC TTCAGCCGCC CGGAGCCGCT CCCGGAGCCC GGCCGTAGAG GCTGCAATCG 120 20 CAGCCGGGAG CCCGCAGCCC GCGCCCCGAG CCCGCCGCCG CCCTTCGAGG GCGCCCCAGG 180 CCGCGCCATG GTGAAGGTGA CGTTCAACTC CGCTCTGGCC CAGAAGGAGG CCAAGAAGGA 240 CGAGCCCAAG AGCGGCGAGG AGGCGCTCAT CATCCCCCCC GACGCCGTCG CGGTGGACTG 300 25 CAAGGACCCA GATGATGTGG TACCAGTTGG CCAAAGAAGA GCCTGGTGTT GGTGCATGTG 360 CTTTGGACTA GCATTTATGC TTGCAGGTGT TATTCTAGGA GGAGCATACT TGTACAAATA 420 30 TTTTGCACTT CAACCAGATG ACGTGTACTA CTGTGGAATA AAGTACATCA AAGATGATGT 480 CATCTTAAAT GAGCCCTCTG CAGATGCCCC AGCTGCTCTC TACCAGACAA TTGAAGAAAA 540 TATTAAAATC TTTGAAGAAG AAGAAGTTGA ATTTATCAGT GTGCCTGTCC CAGAGTTTGC 600 35 AGATAGTGAT CCTGCCAACA TTGTTCATGA CTTTAACAAG AAACTTACAG CCTATTTAGA 660 TCTTAACCTG GATAAGTGCT ATGTGATCCC TCTGAACACT TCCATTGTTA TGCCACCCAG 720 40 AAACCTACTG GAGTTACTTA TTAACATCAA GGCTGGAACC TATTTGCCTC AGTCCTATCT 780 GATTCATGAG CACATGGTTA TTACTGATCG CATTGAAAAC ATTGATCACC TGGGTTTCTT 840 TATTTATCGA CTGTGTCATG ACAAGGAAAC TTACAAACTG CAACGCAGAG AAACTATTAA 900 45 AGGTATTCAG AAACGTGAAG CCAGCAATTG TTTCGCAATT CGGCATTTTG AAAACAAATT 960 TGCCGTGGAA ACTITAATTT GTTCTTGAAC AGTCAAGAAA AACATTATTG AGGAAAATTA 1020 50 ATATCACAGC ATAACCCCAC CCTTTACATT TTGTGCAGTG ATTATTTTTT AAAGTCTTCT 1080 TTCATGTAAG TAGCAAACAG GGCTTTACTA TCTTTTCATC TCATTAATTC AATTAAAACC 1140 ATTACCTTAA AATTTTTTC TTTCGAAGTG TGGTGTCTTT TATATTTGAA TTAGTAACTG 1200 55 TATGAAGTCA TAGATAATAG TACATGTCAC CTTAGGTAGT AGGAAGAATT ACAATTTCTT 1260 TAAATCATTT ATCTGGATTT TTATGTTTTA TTAGCATTTT CAAGAAGACG GATTATCTAG 1320 60 AGAATAATCA TATATATGCA TACGTAAAAA TGGACCACAG TGACTTATTT GTAGTTGTTA 1380

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	GTTGCCCTGC TACCTAGTTT GTTAGTGCAT TTGAGCACAC ATTTTAATTT TCCTCTAATT	1440
ہے	AAAATGTGCA GTATTTTCAG TGTCAAATAT ATTTAACTAT TTAGAGAATG ATTTCCACCT	1500
5	TTATGTTTTA ATATCCTAGG CATCTGCTGT AATAATATTT TAGAAAATGT TTGGAATTTA	1560
	AGAAATAACT TGTGTTACTA ATTTGTATAA CCCATATCTG TGCAATGGAA TATAAATATC	1620
10	ACAAAGTTGT TTAAMWAAAA AAAAAAAAA AAAAAAAAA AAAAAAAAA AAAAA	1679
15	(2) INFORMATION FOR SEQ ID NO: 120:	
	(i) SEQUENCE CHARACTERISTICS:	
20	(A) LENGTH: 1308 base pairs(B) TYPE: nucleic acid(C) STRANDEDNESS: double(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 120:	
25	TTGGCANCNG GGAGAGGGAA AGAGGAGGAA ATGGGGTTTG AGGACCATGG CTTACCTTTC	60
	CTGCCTTTGA CCCATCACAC CCCATTTCCT CCTCTTTCCC TCTCCCCGCT GCCAAAAAAA	120
	AAAAAAAGG AAACGTTTAT CATGAATCAA CAGGGTTTCA GTCCTTATCA AAGAGAGATG	180
30	TGGAAAGAGC TAAAGAAACC ACCCTTTGTT CCCAACTCCA CTTTACCCAT ATTTTATGCA	240
	ACACAAACAC TGTCCTTTTG GGTCCCTTTC TTACAGATGG ACCTCTTGAG AAGAATTATC	300
35	GTATTCCACG TTTTTAGCCC TCAGGTTACC AAGATAAATA TATGTATATA TAACCTTTAT	360
	TATTGCTATA TCTTTGTGGA TAATACATTC AGGTGGTGCT GGGTGATTTA TTATAATCTG	420
40	AACCTAGGTA TATCCTTTGG TCTTCCACAG TCATGTTGAG GTGGGCTCCC TGGTATGGTA	480
40	AAAAGCCAGG TATAATGTAA CTTCACCCCA GCCTTTGTAC TAAGCTCTTG ATAGTGGATA	540
	TACTCTTTTA AGTTTAGCCC CAATATAGGG TAATGGAAAT TTCCTGCCCT CTGGGTTCCC	600
45	CATTTTTACT ATTAAGAAGA CCAGTGATAA TTTAATAATG CCACCAACTC TGGCTTAGTT	660
	AAGTGAGAGT GTGAACTGTG TGGCAAGAGA GCCTCACACC TCACTAGGTG CAGAGAGCCC	720
50	AGGCCTTATG TTAAAATCAT GCACTTGAAA AGCAAACCTT AATCTGCAAA GACAGCAGCA	780
30	AGCATTATAC GGTCATCTTG AATGATCCCT TTGAAATTTT TTTTTTGTTT GTTTGTTTAA	840
	ATCAAGCCTG AGGCTGGTGA ACAGTAGCTA CACACCCATA TTGTGTGTTC TGTGAATGCT	900
55	AGCTCTCTTG AATTTGGATA TTGGTTATTT TTTATAGAGT GTAAACCAAG TTTTATATTC	960
	TGCAATGCGA ACAGGTACCT ATCTGTTTCT AAATAAAACT GTTTACATTC ATTATGGGGT	1020
	ATGTATGACC TTCATTTTCC AAGAAATAGA ACTCTAGCTT AGAATTATGG ATGCTCTAAA	1080

	ATGTCAGAAT	GGGAACTCTC	CTCGAAGTTC	TCCCAAACTC	AGAGACAGCA	CTGCCTTCTC	1140
	CTAAATGATT	ATTCTTTTCT	CCCTGTTTTC	TGGTATTTTC	TAGGCATCCT	TCTCACCACA	1200
5	GCCATAACCC	тттттастт	CCATTAGGCC	GTATAACTGG	NGGGACNGCT	GGTCGGTATA	1260
	TAATACTGGT	WCCAACAMAG	GGGTTCTGGA	TGTACACMAG	GTTATCTT		1308
10							
10	(2) 7770710	m					
			EQ ID NO: 1				
15	(1)	(A) LEN (B) TYP (C) STR	HARACTERIST GTH: 1411 b E: nucleic ANDEDNESS: OLOGY: line	ase pairs acid double			
20	(xi)	SEQUENCE 1	DESCRIPTION	: SEQ ID NO	: 121:		
	GGCACAGGAG	CGACCCGGGA	GAAGGAGGC	CAMGAKGCGG	AAGCGGAGGA	GTCTCCAGGA	60
25	GACCCGGGGA	CAGCATCGCC	CAGGCCCCTG	TTTGCAGGCC	TTTCAGATAT	ATCCATCTCA	120
	CAAGACATCC	CCGTAGAAGG	AGAAATCACC	ATTCCTATGA	GATCTCGCAT	CCGGGAGTTT	180
	GACAGCTCCA	CATTAAATGA	ATCTGTTCGC	AATACCATCA	TGCGTGATCT	AAAAGCTGTT	240
30	GGGAAAAAAT	TCATGCATGT	TTTGTACCCA	AGGAAAAGTA	ATACTCTTTT	GAGAGATTGG	300
	GATTTGTGGG	GCCCTTTGAT	CCTTTGTGTG	ACACTCGCAT	TAATGCTGCA	AAGAGACTCT	360
35	GCAGATAGTG	AAAAAGATGG	AGGGCCCCAA	TTTGCAGAGG	TGTTTGTCAT	TGTCTGGTTT	420
55	GGTGCAGTTA	CCATCACCCT	CAACTCAAAA	CTTCTTGGAG	GGAACATATC	TTTTTTCAG	480
	AGCCTCTGTG	TGCTGGGTTA	CTGTATACTT	CCCTTGACAG	TAGCAATGCT	GATTTGCCGG	540
40	CTGGTACTTT	TGGCTGATCC	AGGACCTGTA	AACTTCATGG	TTCGGCTTTT	TGTGGTGATT	600
	GTGATGTTTG	CCTGGTCTAT	AGTTGCCTCC	ACAGCTTTCC	TTGCTGATAG	CCAGCCTCCA	660
45	AACCGCAGAG	CCCTAGCTGT	TTATCCTGTT	TTCCTGTTTT	ACTITGTCAT	CAGTTGGATG	720
	ATTCTCACCT	TTACTCCTCA	GTAAATCAGG	AATGGGAAAT	TAAAAACCAG	TGAATTGAAA	780
	GCACATCTGA	AAGATGCAAT	TCACCATGGA	GCTTTGTCTC	TGGCCCTTAT	TTGTCTAATT	840
50	TTGGAGGTAT	TIGATAACTG	AGTAGGTGAG	GAGATTAAAA	GGGAGCCATA	TAGCACTGTC	900
	ACCCCTTATT	TGAGGAACTG	ATGTTTGAAA	GGCTGTTCTT	TTCTCTCTTA	ATGTCATTTC	960
55	TTTAAAAATA	CATGTGCATA	CTACACACAG	TATATAATGC	CTCCTTAAGG	CATGATGGAG	1020
5 5	TCACCGTGGT	CCATTTGGGT	GACAACCAGT	GACTTGGGAA	GCACATAGAT	ACATCTTACA	1080
	AGTTGAATAG	AGTTGATAAC	TATTTTCAGT	TTTGAGAATA	CCAGTTCAGG	TGCAGCTCTT	1140
60	AAACACATTG	CCTTATGACT	ATTAGAATAT	GCCTCTCTTT	тсатааатаа	AAATACATGG	1200

	TCTATATCCA TTTTCTTTTA TTTCTCTCTC TTAAGCTTAA AAAGGCAATG AGAGAGGTTA	1260
_	GGAGTGGGTT CATACACGGA GAATGAGAAA ACATGCATTA ACCAATATTC AGATTTTGAT	1320
5	CAGGGGAAAT TCTAYACTTG TTGCAAAAAA AAAAAAAAA AAACTCGAGG GGGGCCCGGT	1380
	ACCCAATCGC NGTATATGAT CGNAAACAAT C	1411
10		
	(2) INFORMATION FOR SEQ ID NO: 122:	
15	(i) SEQUENCE CHARACTERISTICS:	
15	(A) LENGTH: 2256 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear	
20	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 122:	
	GCTTTGGCTT TTTTTGGCGG ACTGGGGCGC CCTCCGGAAG CGTTTCCAAC TTTCCAGAAG	60
25	TTTCTCGGGA CGGGCAGGAG GGGGTGGGGA CTGCCATATA TAGATCCCGG GAGCAGGGGA	120
	GCGGGCTAAG AGTAGAATCG TGTCGCGGCT CGAGAGCGAG AGTCACGTCC CGGCGCTAGC	180
30	CAGCCCGACC CAGGCCCACC GTGGTGCACG CAAACCACTT CCTGGCCATG CGCTCCCTCC	240
30	TGCTTCTCAG CGCCTTCTGC CTCCTGGAGG CGGCCCTGGC CGCCGAGGTG AAGAAACCTG	300
	CAGCCGCAGC AGCTCCTGGC ACTGCGGAGA AGTTGAGCCC CAAGGCGGCC ACGCTTGCCG	360
35	AGCGCANGCC GGCCTGGCCT TCAGCTTGTA CCAGGCCATG GCCAAGGACC AGGCAGTGGA	420
	GAACATCCTG GTGTCACCCG TGGTGGTGGC CTCGTCGCTG GGGCTCGTGT CGCTGGGCGG	480
40	CAAGGCGACC ACGGCGTCGC AGGCCAAGGC AGTGCTGAGC GCCGAGCAGC TGCGCGACGA	540
40	GGAGGTGCAC GCCGGCCTGG GCGAGCTGCT GCGCTCACTC AGCAACTCGA CGGCGCGCAA	600
	CGTGACCTGG AAGCTGGGCA GCCGACTGTA CGGACCCAGC TCAGTGAGCT TCGCTGATGA	660
45	CTTCGTGCGC ACAGCAAGCA GCACTACAAC TGCGAGCACT CCAAGATCAA CTTCCGCGAC	720
	AAGCGCAGNG CGCTGCAGTC CATCAACGAG TGGGCCGCGC AGACCACCGA CGGCAAGCTG	780
50	CCCGAGGTCA CCAAGGACGT GGAGCGCACG GACGGCGCCC TGCTAGTCAA CGCCATGTTC	840
30	TTCAAGCCAC ACTGGGATGA GAAATTCCAC CACAAGATGG TGGACAACCG TGGCTTCATG	900
	GTGACTCGGT CCTATACYGT GGGTGTCATG ATGATGCACC GGACAGGCCT CTACAACTAC	960
55	TACGACGACG AGAAGGAAAA GCTGCAAATC GTGGAGATGC CCCTGGCCCA CAAGCTCTCC	1020
	AGCCTCATCA TCCTCATGCC CCATCACGTG GAGCCTCTCG AGCGCCTTGA AAAGCTGCTA	1080
60	ACCAAAGAGC AGCTGAAGAT CTGGATGGGG AAGATGCAGA AGAAGGCTGT TGCCATCTCC	1140
00		

	TTGCCCAAGG	GTGTGGTGGA	GGTGACCCAT	GACCTGCAGA	AACACCTGGC	TGGGCTGGGC	1200
	CTGACTGAGG	CCATTGACAA	GAACAAGGCC	GACTTRTCAC	GCATGTCAGG	CAAGAAGGAC	1260
5	CTGTACCTGG	CCAGCGTGTT	CCACGCCACC	GCCTTTGAGT	TGGACACAGA	TGGCAACCCC	1320
	TTTGACCAGG	ACATCTACGG	GCGCGAGGAG	CTGCGCANCC	CAAGCTGTTC	TACGCCGACC	1380
10	ACCCCTTCAT	CTTCCTAGTG	CGGGACACCC	AAAGCGGCTC	CCTGCTATTC	ATTGGGCGCC	1440
•	TGGTCCGGCC	TAAGGGTGAC	AAGATGCGAG	ACGAGTTATA	GGCCTCAGG	GTGCACACAG	1500
	GATGGCAGGA	GGCATCCAAA	GGCTCCTGAG	ACACATGGGT	GCTATTGGGG	TTGGGGGGA	1560
15	GGTGAGGTAC	CAGCCTTGGA	TACTCCATGG	GGTGGGGGTG	GAAAARCAGA	CCGGGGTTCC	1620
	CGTGTGCCTG	AGCGGACCTT	CCCAGCTAGA	ATTCACTCCA	CTTGGACATG	GGCCCCAGAT	1680
20	ACCATGATGC	TGAGCCCGGA	AACTCCACAT	CCTGTGGGAC	CTGGGCCATA	GTCATTCTGC	1740
20	CTGCCCTGAA	AGTCCCAGAT	CAAGCCTGCC	TCAATCAGTA	TTCATATTTA	TAGCCAGGTA	1800
	CCTTCTCACC	TGTGAGACCA	AATTGAGCTA	GGGGGTCAG	CCAGCCCTCT	TCTGACACTA	1860
25	AAACACCTCA	GCTGCCTCCC	CAGCTCTATC	CCAACCTCTC	CCAACTATAA	AACTAGGTGC	1920
	TGCAGCCCCT	GGGACCAGGC	ACCCCCAGAA	TGACCTGGCC	GCAGTGAGGC	GGATTGAGAA	1980
30	GGAGCTCCCA	GGAGGGGCTT	CTGGGCAGAC	TCTGGTCAAG	AAGCATCGTG	TCTGGCGTTG	2040
	TGGGGATGAA	CTTTTTGTTT	TGTTTCTTCC	TTTTTTAGTT	CTTCAAAGAT	AGGGAGGGAA	2100
	GGGGAACAT	GAGCCTTTGT	TGCTATCAAT	CCAAGAACTT	ATTTGTACAT	TTTTTTTTC	2160
35	AATAAAACTT	TTCCAATGAC	AAAAAAAA	AAAAAAAA	AAAAAGGGGS	GGGCCGCTCC	2220
	TAGAGGGATC	CCTCCGANGG	NGCCCAATCG	NTAAAA			2256
40							
	(2) INFORMA	TION FOR SE	Q ID NO: 12)			
45		SEQUENCE CH (A) LENG (B) TYPI (C) STR	HARACTERIST: GTH: 829 bases: nucleic and ANDEDNESS: OLOGY: line	ICS: se pairs acid double			
50	(xi)	SEQUENCE I	DESCRIPTION	SEQ ID NO:	: 123:		
	ATGCGCTCCC	TCCTGCTTCT	CAGCGCCTTC	TGCCTCCTGG	AGGCGGCCCT	GGCCGCCGAG	60
55	GTGAAGAAAC	CTGCAGCCGC	AGCAGCTCCT	GGCACTGCGG	AGAAGTTGAG	CCCCAAGGCG	120
,,,	GCCACGCTTG	CCGAGCGCAA	GCGGCCTGGC	CTTCAGCTIG	TACCAGGCCA	TGGCCAAGGA	180
	CCAGGCAGTG	GAGAACATCC	TGGTGTCACC	CGTGGTGGTG	GCCTCGTCGC	TGGGGCTCGT	240
60	GTCGCTGGGC	GGCAAGGCGA	CCACGGCGTC	GCAGGCCAAG	GCAGTGCTGA	GCGCCGAGCA	300

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	GCTGCGCGAC GAGGAGGTGC ACGCCGGCCT GGGCGAGCTG CTGCGCTCAC TCAGCAACTC	360
	CACGGCGCGC AACGTGACCT GGAAGCTGGG CAGCCGACTG TACGGACCCA GCTCAGTGAG	420
5	CTTCGCTGAT GACTTCGTGC GCAGCAGCAA GCAGCACTAC AACTGCGAGC ACTCCAAGAT	480
	CAACTICCGC GACAAGCGCA GCGCGCTGCA GTCCATCAAC GAGTGGGCCG CGCAGACCAC	540
10	CGACGGCAAG CTGCCCGAGG TCACCAAGGA CGTGGAGGGC ACGGACGGCG CCCTGTTAGT	600
	CAACGCCATG TTCTTCAAGC CACACTGGGA TGAGAAATTC CACCACAAGA TGGTGGACAA	660
	CCGTGGCTTC ATGGTGACTC GGTCCTATAC CGTGGGTGTC ATGATGATGC ACCGGACAGG	720
15	CCTCTACAAC TACTACGACG ACGAGAAGGA AAAGCTGCAA ATCGTGGAGA TGCCCCTGGC	780
	CCACAAGCTC TCCAGCCTCA TCATCCTCAT GCCCCATCAC GTGGAGCCT	829
20		
	(2) INFORMATION FOR SEQ ID NO: 124:	
25	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 2223 base pairs	
	(B) TYPE: nucleic acid (C) STRANDEDNESS: double	
30	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 124:	CO
	CCTCCGGAAG CGTTTCCAAC TTTCCAGAAG TTTCTCGGGA CGGGCAGGAG GGGGTGGGGA	60
35	CTGCCATATA TAGATCCCGG GAGCAGGGGA GCGGGCTAAG AGTAGAATCG TGTCGCGGCT	120
	CGAGAGCGAG AGTCACGTCC CGGCGCTAGC CAGCCCGACC CAGGCCCACC GTGGTGCACG	180
40	CAAACCACTT CCTGGCCATG CGCTCCCTCC TGCTTCTCAG CGCCTTCTGC CTCCTGGAGG	240
40	CGGCCCTGGC CGCCGAGGTG AAGAAACCTG CAGCCGCAGC AGCTCCTGGC ACTGCGGAGA	300
	AGTTGAGCCC CAAGGCGGCC ACGCTTGCCG AGCGCAGNCG GCCTGGCCTT CAGCTTGTAC	360
45	CAGGCCATGG CCAAGGACCA GGCAGTGGAG AACATCCTGG TGTCACCCGT GGTGGTGGCC	420
	TCGTCGCTGG GGCTCGTGTC GCTGGGCGGC AAGGCGACCA CGGCGTCGCA GGCCAAGGCA	480
50	GTGCTGAGCG CCGAGCAGCT GCGCGACGAG GAGGTGCACG CCGGCCTGGG CGAGCTGCTG	540
50	CGCTCACTCA GCAACTCSAC GGCGCGCAAC GTGACCTGGA AGCTGGGCAG CCGACTGTAC	600
	GGACCCAGCT CAGTGAGCTT CGCTGATGAC TTCGTGCGCA CAGCAAGCAG CACTACAACT	660
55	GCGAGCACTC CAAGATCAAC TTCCGCGACA AGCGCACGCG CTGCAGTCCA TCAACGAGTG	720
	GGCCGCGCAG ACCACCGACG GCAAGCTGCC CGAGGTCACC AAGGACGTGG AGCGCACGGA	780
	CGGCGCCCTG YTAGTCAACG CCATGTTCTT CAAGCCACAC TGGGATGAGA AATTCCACCA	840

840

	CAAGATGGTG	GACAACCGTG	GCTTCATGGT	GACTCGGTCC	TATACYGTGG	GTGTCATGAT	900
	GATGCACCGG	ACAGGCCTCT	ACAACTACTA	CGACGACGAG	AAGGAAAAGC	TGCAAATCGT	960
5	GGAGATGCCC	CTGGCCCACA	AGCTCTCCAG	CCTCATCATC	CTCATGCCCC	ATCACGTGGA	1020
	GCCTCTCGAG	CGCCTTGAAA	AGCTGCTAAC	CAAAGAGCAG	CTGAAGATCT	GGATGGGGAA	1080
10	GATGCAGAAG	AAGGCTGTTG	CCATCTCCTT	GCCCAAGGGT	GTGGTGGAGG	TGACCCATGA	1140
	CCTGCAGAAA	CACCTGGCTG	GGCTGGGCCT	GACTGAGGCC	ATTGACAAGA	ACAAGGCCGA	1200
	CTTRTCACGC	ATGTCAGGCA	AGAAGGACCT	GTACCTGGCC	AGCGTGTTCC	ACGCCACCGC	1260
15	CTTTGAGTTG	GACACAGATG	GCAACCCCTT	TGACCAGGAC	ATCTACGGGC	GCGAGGAGCT	1320
	GCGCASCCCA	AGCTGTTCTA	CGCCGACCAC	CCCTTCATCT	TCCTAGTGCG	GGACACCCAA	1380
20	AGCGGCTCCC	TGCTATTCAT	TGGGCGCCTG	GTCCGGCCTA	AGGGTGACAA	GATGCGAGAC	1440
	GAGTTATAGG	GCCTCAGGGT	GCACACAGGA	TGGCAGGAGG	CATCCAAAGG	CTCCTGAGAC	1500
	ACATGGGTGC	TATTGGGGTT	GGGGGGAGG	TGAGGTACCA	GCCTTGGATA	CTCCATGGGG	1560
25	TGGGGGTGGA	AAARCAGACC	GGGGTTCCCG	TGTGCCTGAG	CGGACCTTCC	CAGCTAGAAT	1620
	TCACTCCACT	TGGACATGGG	CCCCAGATAC	CATGATGCTG	AGCCCGGAAA	CTCCACATCC	1680
30	TGTGGGACCT	GGGCCATAGT	CATTCTGCCT	GCCCTGAAAG	TCCCAGATCA	AGCCTGCCTC	1740
	AATCAGTATT	CATATTTATA	GCCAGGTACC	TTCTCACCTG	TGAGACCAAA	TTGAGCTAGG	1800
	GGGGTCAGCC	AGCCCTCTTC	TGACACTAAA	ACACCTCAGC	TGCCTCCCCA	GCTCTATCCC	1860
35	AACCTCTCCC	AACTATAAAA	CTAGGTGCTG	CAGCCCCTGG	GACCAGGCAC	CCCCAGAATG	1920
	ACCTGGCCGC	AGTGAGGCGG	ATTGAGAAGG	AGCTCCCAGG	AGGGGCTTCT	GGGCAGACTC	1980
40	TGGTCAAGAA	GCATCGTGTC	TGGCGTTGTG	GGGATGAACT	TTTTGTTTTG	TTTCTTCCTT	2040
	TTTTAGTTCT	TCAAAGATAG	GGAGGGAAGG	GGGAACATGA	GCCTTTGTTG	CTATCAATCC	2100
	AAGAACTTAT	TTGTACATTT	TTTTTTCAA	TAAAACTTTT	CCAATGACAA	AAAAAAAA	2160
45	AAAAAAAA	MWMGGGGSGG	GCCGCTCCTA	GAGGGATCCC	TCCGANGGNG	CCCAATCGAA	2220
	TAA						2223

55

(2) INFORMATION FOR SEQ ID NO: 125:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 31 amino acids

(B) TYPE: amino acid

(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 125:

 $60\,$ Met Lys Lys Gln Ser Lys Arg Cys Leu Trp Lys Pro Pro Gly Ser Leu

	1				5					10					15	
5	Arg	Arg	Leu	Trp 20	Trp	Met	Arg	Ala	Leu 25	Leu	Ile	Leu	Lys	Туr 30	Ile	
5																
	(2)	INF	ORMA	rion	FOR	SEQ	ID 1	10: 1	L26:							
10			(i)	(A) L B) T	ENGT YPE :	H: 4 ami		ino a		s					
15			(xi)	SEQ!	UENC:	E DE	SCRI	PTIO	N: SI	EQ I	D NO	: 12	6:			
	Met 1	Lys	Lys	Ser	Leu 5	Glu	Asn	Leu	Asn	Arg 10	Leu	Gln	Val	Met	Leu 15	Leu
20	His	Leu	Thr	Ala 20	Ala	Phe	Leu	Gln	Arg 25	Ala	His	Xaa	Ile	Leu 30	Thr	Thr
	Arg	Met	Ser 35	Leu	Gly	Phe	Gln	Ser 40	Pro	His	Leu	Thr	Met 45			
25																
	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	127 :							
30			, ,		(A) I (B) T (D) T	ENGT TYPE : TOPOI	H: 3 ami OGY:	9 an .no a : lir	nino acid near	acid): 12	7:			
35	Met		: Asr	ı Gln	Arg 5		Val	Phe	. Leu	Phe 10		Leu	Phe	Ser	Asn 15	Tyr
40				20 20 Met					Gly 25		Leu	Leu	Ala	. Ala 30		Tyr
45	(2)	IN	FORM	AOITA	1 FOF	R SEÇ) ID	NO:	128:			-				
50				SEQI	(A) (B) (D)	LENG TYPE TOPO	TH: : am LOGY	23 a ino : li	mino acid near	aci		O: 1	28:			
55		t Ar	g Ly	s Ly:		e Lev	ı Lei	ı Ala	a Glr	1 Va:		e Le	ı Sei	r Lei	ı Ser 15	Val
	Me	t Pr	o Se	r Me		o Vai	l Th	r								
60																

	(2)	INF	ORMA'	TION	FOR	SEQ	ID I	O:	129:							
5			(i) (xi)	(A) L B) T D) T	ENGT YPE : OPOL	RACT H: 1 ami OGY: SCRI	10 a no a lin	mino cid ear	aci		: 12	9:			
10	Met 1	Val	Leu	Leu	Cys 5	Leu	Leu	Leu	Val	Pro 10	Leu	Leu	Leu	Ser	Leu 15	Phe
15	Val	Leu	Gly	Leu 20	Phe	Leu	Trp	Phe	Leu 25	Lys	Arg	Glu	Arg	Gln 30	Glu	Glu
	Tyr	Ile	Glu 35	Glu	Lys	Lys	Arg	Val 40	Asp	Ile	Cys	Arg	Glu 45	Thr	Pro	Asn
20	Ile	Cys 50	Pro	His	Ser	Gly	Glu 55	Asn	Thr	Glu	Tyr	Asp 60	Thr	Ile	Pro	His
	Thr 65	Asn	Arg	Thr	Ile	Leu 70	Lys	Glu	Asp	Pro	Ala 75	Asn	Thr	Val	Tyr	Ser 80
25	Thr	Val	Glu	Ile	Pro 85	Lys	Lys	Met	Glu	Asn 90	Pro	His	Ser	Leu	Leu 95	Thr
30	Met	Pro	Asp	Thr 100	Pro	Arg	Leu	Phe	Ala 105	Tyr	Glu	Asn	Val	Ile 110		
	(2)	INF	ORMA'	rion	FOR	SEQ	ID 1	NO: 1	L30:	•						
35				(A) L B) T D) T	ENGT YPE : OPOL	H: 6 ami OGY:	3 am no a lin	ino cid ear	acid	•	. 12	n			
40	Mo≠	Lou		SEQ										_		_
	1		Leu		5	,				10					15	
15	Ala	Gly	Ala	Thr 20	Ser	Lys	Pro	Arg	Tyr 25	Arg	Val	Ile	Thr	Суs 30	Gly	Pro
	Ala	Ser	Val 35	Phe	Ser	Thr	Ser	Phe 40	Ser	His	Ser	Pro	Pro 45	Ala	Arg	Cys
50	Leu	Gly 50	Arg	Leu	Glu	Gln	Met 55	Phe	His	Phe	Gly	Leu 60	Ala	Ser	Gly	
55	(2)	INFO	ORMA	SEQU:	ENCE	CHAI		ERIS	rics		s					
60							ami: OGY:									

```
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 131:
     Met Pro Phe Pro Ile Ser Ile Leu Gln Leu Cys Leu Gln Ile Ser Asn
       1
 5
     Leu Ser Phe Cys Leu Gln Lys Ile Tyr Lys Ile Pro Phe Val
                                      25
10
     (2) INFORMATION FOR SEQ ID NO: 132:
             (i) SEQUENCE CHARACTERISTICS:
                   (A) LENGTH: 53 amino acids
15
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 132:
      Met Ala Ala Ala Cys Arg Ser Val Lys Gly Leu Val Ala Val Ile Thr
20
      Gly Gly Ala Ser Gly Leu Gly Leu Ala Thr Ala Asp Asp Leu Trp Gly
                                      25
      Arg Glu Pro Leu Cys Phe Trp Thr Cys Pro Thr Arg Val Gly Arg
25
                                 40
              35
      Pro Lys Pro Arg Ser
          50
30
      (2) INFORMATION FOR SEQ ID NO: 133:
35
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 57 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 133:
40
      Met Leu Leu Val Tyr Asp Leu Tyr Leu Xaa Pro Lys Leu Trp Ala Leu
                                           10
      Ala Thr Pro Gln Lys Asn Gly Lys Gly Ala Arg Xaa Gly Asp Gly Thr
45
                   20
      Pro Ala Gln Ala Phe Trp Asp Phe Trp Ser His Leu Ile Ser Ala Asp
50
      Pro Gln Thr Trp Glu Arg Ala Ala Pro
                             55
55
      (2) INFORMATION FOR SEQ ID NO: 134:
              (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 216 amino acids
                     (B) TYPE: amino acid
 60
                     (D) TOPOLOGY: linear
```

			(xi)	SEQ	UENCI	E DE	SCRI	PTIO	N: S	EQ II	ои с	: 13	4:			
5	Met 1	Arg	Leu	Ser	Ala 5	Leu	Leu	Ala	Leu	Ala 10	Ser	Lys	Val	Thr	Leu 15	Pro
J	Pro	His	Tyr	Arg 20	Tyr	Gly	Met	Ser	Pro 25	Pro	Gly	Ser	Val	Ala 30	Asp	Lys
10	Arg	Lys	Asn 35	Pro	Pro	Trp	Ile	Arg 40	Arg	Arg	Pro	Val	Val 45	Val	Glu	Pro
	Ile	Ser 50	Asp	Glu	Asp	Trp	Tyr 55	Leu	Phe	Cys	Gly	Asp 60	Thr	Val	Glu	Ile
15	Leu 65	Glu	Gly	Lys	Asp	Ala 70	Gly	Lys	Gln	Gly	Lys 75	Val	Val	Gln	Val	Ile 80
20	Arg	Gln	Arg	Asn	Trp 85	Val	Val	Val	Gly	Gly 90	Leu	Asn	Thr	His	Туг 95	Arg
	Tyr	Ile	Gly	Lys 100	Thr	Met	Asp	Tyr	Arg 105	Gly	Thr	Met	Ile	Pro 110	Ser	Glu
25	Ala	Pro	Leu 115	Leu	His	Arg	Gln	Val 120	Lys	Leu	Val	Asp	Pro 125	Met	Asp	Arg
	Lys	Pro 130	Thr	Glu	Ile	Glu	Trp 135	Arg	Phe	Thr	Glu	Ala 140	Gly	Glu	Arg	Val
30	Arg 145	Val	Ser	Thr	Arg	Ser 150	Gly	Arg	Ile	Ile	Pro 155	Lys	Pro	Glu	Phe	Pro 160
35	Arg	Ala	Asp	Gly	Ile 165	Val	Pro	Glu	Thr	Trp 170	Ile	Asp	Gly	Pro	Lys 175	Asp
	Thr	Ser	Val	Glu 180	Asp	Ala	Leu	Glu	Arg 185	Thr	Tyr	Val	Pro	Cys 190	Leu	Lys
40	Thr	Leu	Gln 195	Glu	Glu	Val	Met	Glu 200	Ala	Met	Gly	Ile	Lys 205	Glu	Thr	Arg
	Lys	Tyr 210	Lys	Lys	Val	Tyr	Trp 215	Tyr								
45	(2)	INF	orma'	TION	FOR	SEQ	ID :	NO:	135:							
50				(ENCE (A) I (B) I (D) I	ENGT YPE : OPOL	H: 4 ami OGY:	19 am no a lin	ino cid ear	acid): 1 3	5:			
55	Met 1		Leu	Arg	Gln 5	Lys	Ser	Ser	Phe	Arg 10	Leu	Met	Val	Met	Ser 15	Let
60	Thr	Ile	. Leu	Lys 20		Ser	Lys	Thr	Thr 25		Leu	Суз	Leu	Arg 30	Cys	Leu

```
His Ser Leu Lys Leu Thr Trp Arg Asp Gly Ala Arg Cys Ile Asn Ala
                                   40
     Glu
 5
      (2) INFORMATION FOR SEQ ID NO: 136:
10
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 68 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
15
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 136:
     Met Ser Gly Ser Phe Ile Leu Cys Leu Ala Leu Val Thr Arg Trp Ser
20
      Pro Gln Ala Ser Ser Val Pro Leu Ala Val Tyr Glu Ser Lys Thr Arg
                                       25
      Lys Ser Tyr Arg Ser Gln Arg Asp Arg Asp Gly Lys Asp Arg Ser Gln
                                  40
25
      Gly Met Gly Leu Ser Leu Leu Val Glu Thr Arg Lys Leu Leu Leu Ser
                              55
      Ala Asn Gln Gly
30
      65
      (2) INFORMATION FOR SEQ ID NO: 137:
35
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 52 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
40
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 137:
      Met Cys Phe Arg Phe Phe Leu Phe Cys Ser Arg Ile Leu Leu Lys Leu
                       5
45
      Phe Phe Leu Leu Phe Pro Ala Ser Ala Phe Pro Leu Ser Thr Arg Ser
                                       25
      Ser Leu Ser Val Asn Glu His Val Val Val Ser Pro Arg Ser Thr Val
                                   40
                                                      45
50
      Ser Ile Ser Arg
          50
55
      (2) INFORMATION FOR SEQ ID NO: 138:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 541 amino acids
60
                    (B) TYPE: amino acid
```

280

(D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 138:

5	Met 1	Val	Arg	Thr	Asp 5	Gly	His	Thr	Leu	Ser 10	Glu	Lys	Arg	Asn	Tyr 15	Gln
	Val	Thr	Asn	Ser 20	Met	Phe	Gly	Ala	Ser 25	Arg	Lys	Lys	Phe	Val 30	Glu	Gly
10	Val	Asp	Ser 35	Asp	Tyr	His	Asp	Glu 40	Asn	Met	Tyr	Tyr	Ser 45	Gln	Ser	Ser
15	Met	Phe 50	Pro	His	Arg	Ser	Glu 55	Lys	Asp	Met	Leu	Ala 60	Ser	Pro	Ser	Thr
13	Ser 65	Gly	Gln	Leu	Ser	Gln 70	Phe	Gly	Ala	Ser	Leu 75	Tyr	Gly	Gln	Gln	Ser 80
20	Ala	Leu	Gly	Leu	Pro 85	Met	Arg	Gly	Met	Ser 90	Asn	Asn	Thr	Pro	Gln 95	Leu
	Asn	Arg	Ser	Leu 100	Ser	Gln	Gly	Thr	Gln 105	Leu	Pro	Ser	His	Val 110	Thr	Pro
25	Thr	Thr	Gly 115	Val	Pro	Thr	Met	Ser 120	Leu	His	Thr	Pro	Pro 125	Ser	Pro	Ser
30	Arg	Gly 130	Ile	Leu	Pro	Met	Asn 135	Pro	Xaa	Asn	Met	Met 140	Asn	His	Ser	Gln
	Val 145	Gly	Gln	Gly	Ile	Gly 150	Ile	Pro	Ser	Arg	Thr 155	Asn	Ser	Met	Ser	Ser 160
35	Ser	Gly	Leu	Gly	Ser 165	Pro	Asn	Arg	Ser	Ser 170	Pro	Ser	Ile	Ile	Cys 175	Met
	Pro	Lys	Gln	Gln 180	Pro	Ser	Arg	Gln	Pro 185	Phe	Thr	Val	Asn	Ser 190	Met	Ser
40	Gly	Phe	Gly 195	Met	Asn	Arg	Asn	Gln 200	Ala	Phe	Gly	Met	Asn 205	Asn	Ser	Leu
45	Ser	Ser 210	Asn	Ile	Phe	Asn	Gly 215	Thr	Asp	Gly	Ser	Glu 220	Asn	Val	Thr	Gly
	L eu 225	Asp	Leu	Ser	Asp	Phe 230	Pro	Ala	Leu	Ala	Asp 235	Arg	Asn	Arg	Arg	Glu 240
50	Gly	Ser	Gly	Asn	Pro 245	Thr	Pro	Leu	Ile	Asn 250	Pro	Leu	Ala	Gly	Arg 255	Ala
	Pro	Tyr	Val	Gly 260	Met	Val	Thr	Lys	Pro 265	Ala	Asn	Glu	Gln	Ser 270	Gln	Asp
5 5	Phe	Ser	Ile 275	His	Asn	Glu	Asp	Phe 280	Pro	Ala	Leu	Pro	Gly 285	Ser	Ser	Tyr
60	Lys	Asp 290	Pro	Thr	Ser	Ser	Asn 295	Asp	Asp	Ser	Lys	Ser 300	Asn	Leu	Asn	Thr

	Ser 305	Gly	Lys	Thr	Thr	Ser 310	Ser	Thŗ	Asp	Gly	Pro 315	Lys	Phe	Pro	Gly	Asp 320
5	Lys	Ser	Ser	Thr	Thr 325	Gln	Asn	Asn	Asn	Gln 330	Gln	Lys	Lys	Gly	Ile 335	Gln
	Val	Leu	Pro	Asp 340	Gly	Arg	Val	Thr	Asn 345	Ile	Pro	Gln	Gly	Met 350	Val	Thr
10	Asp	Gln	Phe 355	Gly	Met	Ile	Gly	Leu 360	Leu	Thr	Phe	Ile	Arg 365	Ala	Ala	Glu
15	Thr	Asp 370	Pro	Gly	Met	Val	His 375	Leu	Ala	Leu	Gly	Ser 380	Asp	Leu	Thr	Thr
	Leu 385	Gly	Leu	Asn	Leu	Asn 390	Ser	Pro	Glu	Asn	Leu 395	Tyr	Pro	Lys	Phe	Ala 400
20	Ser	Pro	Trp	Ala	Ser 405	Ser	Pro	Cys	Arg	Pro 410	Gln	Asp	Ile	Asp	Phe 415	His
	Val	Pro	Ser	Glu 420	Tyr	Leu	Thr	Asn	Ile 4 25	His	Ile	Arg	Asp	Lys 430	Leu	Ala
25	Ala	Ile	Lys 435	Leu	Gly	Arg	Tyr	Gly 440	Glu	Asp	Leu	Leu	Phe 445	Tyr	Leu	Tyr
30	Tyr	Met 450	Asn	Gly	Gly	Asp	Val 455	Leu	Gln	Leu	Leu	Ala 460	Ala	Val	Glu	Leu
	Phe 465	Asn	Arg	Asp	Trp	Arg 470	Tyr	His	Lys	Glu	Glu 475	Arg	Val	Trp	Ile	Thr 480
35	Arg	Ala	Pro	Gly	Met 485	Glu	Pro	Thr	Met	Lys 490	Thr	Asn	Thr	Tyr	Glu 495	Arg
	Gly	Thr	Tyr	Tyr 500	Phe	Phe	Asp	Cys	Leu 505	Asn	Trp	Arg	Lys	Val 510	Ala	Lys
40	Glu	Phe	His 515	Leu	Glu	Tyr	Asp	Lys 520	Leu	Glu	Glu	Arg	Pro 525	His	Leu	Pro
45	Ser	Thr 530	Phe	Asn	Tyr	Asn	Pro 535	Ala	Gln	Gln	Ala	Phe 540	Xaa			
# 0	(2)	INFO	ORMA!	rion	FOR	SEQ	ID I	10:	L39:							
50			(i) :	(A) L B) T	ENGT YPE :	H: 5 ami	ERIS' 8 am no a lin	ino cid		s					
55			(xi)	SEQ	UENC	E DE	SCRI	PTIO	N: S	EQ II	ои о	: 13	9 :			
	Met 1	Ile	Суѕ	Pro	Gln 5	Суѕ	Pro	Leu	Ser	Leu 10	Leu	Cys	Leu	Ile	Ser 15	Ser
60	Leu	Cys	Ser	Leu 20	Val	Ile	Gln	Ile	Ser 25	Leu	Lys	Thr	Ile	Arg 30	Asp	Ile

	Thr	Leu	Leu 35	Asn	Met	Val	Gly	11e 40	Lys	Phe	Ser	Ile	Ser 45	Leu	Ser	Asn
5	Lys	Ile 50	Asn	Ile	Asn	Ser	Arg 55	Thr	Trp	Xaa						
10	(2)	INFO	ORMAT	поп	FOR	SEQ	ID N	NO: 1	L40:							
15			(i) :	(A) L B) T D) T	ENGT YPE : OPOL	H: 2 ami: OGY:	02 a no a lin	mino cid ear	aci		: 14	0:			
20	Met 1	Thr	Leu	Arg	Pro 5	Ser	Leu	Leu	Pro	Leu 10	His	Leu	Leu	Leu	Leu 15	Leu
20	Leu	Leu	Ser	Ala 20	Ala	Val	Cys	Arg	Ala 25	Glu	Ala	Gly	Leu	Glu 30	Thr	Glu
25	Ser	Pro	Val 35	Arg	Thr	Leu	Gln	Val 40	Glu	Thr	Leu	Val	Glu 45	Pro	Pro	Glu
	Pro	Cys 50	Ala	Glu	Pro	Ala	Ala 55	Phe	Gly	Asp	Thr	Leu 60	His	Ile	His	Tyr
30	Thr 65	Gly	Ser	Leu	Val	Asp 70	Gly	Arg	Ile	Ile	Asp 75	Thr	Ser	Leu	Thr	Arg 80
25	Asp	Pro	Leu	Val	Ile 85	Glu	Leu	Gly	Gln	Lys 90	Gln	Val	Ile	Pro	Gly 95	Leu
35	Glu	Gln	Ser	Leu 100	Leu	Asp	Met	Cys	Val 105	Gly	Glu	Lys	Arg	Arg 110	Ala	Ile
40	Ile	Pro	Ser 115	His	Leu	Ala	Tyr	Gly 120	Lys	Arg	Gly	Phe	Pro 125	Pro	Ser	Val
	Pro	Ala 130	Asp	Ala	Val	Val	Gln 135	Tyr	Asp	Val	Glu	Leu 140	Ile	Ala	Leu	Ile
45	Arg 145	Ala	Asn	Tyr	Trp	Leu 150	Lys	Leu	Val	Lys	Gly 155	Ile	Leu	Pro	Leu	Val 160
50	Gly	Met	Ala	Met	Val 165	Pro	Ala	Leu	Leu	Gly 170	Leu	Ile	Gly	Tyr	His 175	Leu
50	Tyr	Arg	Lys	Ala 180	Asn	Arg	Pro	Lys	Val 185	Ser	Lys	Lys	Lys	Leu 190	Lys	Glu
55	Glu	Lys	Arg 195	Asn	Lys	Ser	Lys	Lys 200	Lys	Xaa						
60	(2)	INF	ORMA'	TION	FOR	SEQ	ID 1	NO:	141:							

		1	(i) S	(2		ENGTI	i: 23	17 ar	nino		ds					
5		ı	(xi)	(1	D) TO	OPOLO	ŒΥ:	line	ear	EQ II	ON C	: 141	L:			
	Met 1	Phe	Leu	Arg	Leu 5	Tyr	Leu	Ile	Ala	Arg 10	Val	Met	Leu	Leu	His 15	Ser
10	Lys	Leu	Phe	Thr 20	Asp	Ala	Ser	Ser	Arg 25	Ser	Ile	Gly	Ala	Leu 30	Asn	Lys
15	Ile	Asn	Phe 35	Asn	Thr	Arg	Phe	Val 40	Met	Lys	Thr	Leu	Met 45	Thr	Ile	Cys
15	Pro	Gly 50	Thr	Val	Leu	Leu	Val 55	Phe	Ser	Ile	Ser	Leu 60	Trp	Ile	Ile	Ala
20	Ala 65	Trp	Thr	Val	Arg	Val 70	Cys	Glu	Ser	Pro	Glu 75	Ser	Pro	Ala	Gln	Pro 80
	Ser	Gly	Ser	Ser	Leu 85	Pro	Ala	Trp	Tyr	His 90	Asp	Gln	Gln	Asp	Val 95	Thr
25	Ser	Asn	Phe	Leu 100	Gly	Ala	Met	Trp	Leu 105	Ile	Ser	Ile	Thr	Phe 110	Leu	Ser
20	Ile	Gly	Туг 115	Gly	Asp	Met	Val	Pro 120	His	Thr	Tyr	Cys	Gly 125	Lys	Gly	Val
30	Cys	Leu 130	Leu	Thr	Gly	Ile	Met 135	Gly	Ala	Gly	Cys	Thr 140	Ala	Leu	Val	Val
35	Ala 145		Val	Ala	Arg	Lys 150	Leu	Glu	Leu	Thr	Lys 155		Glu	Lys	His	Val 160
	His	Asn	Phe	Met	Met 165		Thr	Gln	Leu	Thr 170		Arg	Ile	Lys	Asn 175	
40	Ala	Ala	. Asn	Val 180		Arg	Glu	Thr	Trp 185		ılle	Tyr	. Lys	His 190		Lys
45	Leu	Leu	1 Lys 195		: Ile	Asp	His	Ala 200		Val	. Arg	Lys	His 205		. Arg	Lys
45	Phe	210	ı Pro	Ser	туг	Pro	Pro 215		Xaa	ı						
50	(2)	INE	FORM	10ITA	1 FOF	R SEÇ) ID	NO:	142:	:						
55					(A) (B) (D)	LENG TYPE TOPO	TH: : am LOGY	102 ino : li	STIC: amin acid near	o ac		O: 1	42:			
60		t Se: 1	r Ası	n Th		r Val	l Pro	o Ası	n Ala	a Pr		n Ala	a Ası	n Sei	r Ası	

	Met	Val	Gly	Tyr 20		Leu	Gly	Pro	Phe 25	Phe	Leu	Ile	Thr	Leu 30	Val	Gly
5	Val	Val	Val 35	Ala	Val	Val	Met	Tyr 40	Val	Gln	Lys	Lys	Lys 45	Arg	Val	Asp
10	Arg	Leu 50	Arg	His	His	Leu	Leu 55	Pro	Met	Tyr	Ser	Туr 60	Asp	Pro	Ala	Glu
	Glu 65	Leu	His	Glu	Ala	Glu 70	Gln	Glu	Leu	Leu	Ser 75	Asp	Met	Gly	Asp	Pro 80
15	Lys	Val	Val	His	Gly 85	Trp	Gln	Ser	Gly	Туr 90	Gln	His	Lys	Arg	Met 95	Pro
	Leu	Leu	Asp	Val 100	Lys	Thr										
20																
	(2)		ORMA													
25			(i) :						TICS mino		ds					
				(B) T	YPE:	ami	no a lin	cid							
			(xi)							EQ II	OM C	: 14	3 :			
30	Met 1	Arg	Glu	Cys	Gln 5	Glu	Glu	Ser	Phe	Trp 10	Lys	Arg	Ala	Leu	Pro 15	Phe
35	Ser	Leu	Val	Ser 20	Met	Leu	Val	Thr	Gln 25	ГУ	Leu	Val	Tyr	Gln 30	Gly	Tyr
	Leu	Ala	Ala 35	Asn	Ser	Arg	Phe	Gly 40	Sei	Ju	Pro	Lys	Val 45	Ala	Leu	Ala
40	Gly	Leu 50	Leu	Gly	Phe	Gly	Leu 55	Gly	Lys	Val	Ser	Tyr 60	Ile	Gly	Val	Cys
	Gln 65	Ser	Lys	Phe	His	Phe 70	Phe	Glu	Asp	Gln	Leu 75	Arg	Gly	Ala	Gly	Phe 80
45	Gly	Pro	Gln	His	Asn 85	Arg	His	Cys	Leu	Leu 90	Thr	Cys	Glu	Glu	Cys 95	Lys
50	Ile	Lys	His	Gly 100	Leu	Ser	Glu	Lys	Gly 105	Asp	Ser	Gln	Pro	Ser 110	Ala	Ser
55	(2)	INFO	RMAT	CION	FOR	SEQ	ID N	10:1	.44:							
			(i) S						rics:		_					
60								o am no a		aC1ds	5					

```
(D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 144:
     Met Lys Asn Asp Arg Asn Gln Gly Phe Ser Leu Leu Gln Leu Ile Asp
5
                                          10
     Trp Asn Lys Pro
10
      (2) INFORMATION FOR SEQ ID NO: 145:
             (i) SEQUENCE CHARACTERISTICS:
15
                    (A) LENGTH: 30 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 145:
     Met Gly Thr Gln Pro Pro Val Val Ala Gly Phe Thr Ile Pro Met Leu
20
      Gly Tyr Thr Val Arg Val Leu Thr Phe His Leu Ser Cys Ser
                  20
                                      25
25
      (2) INFORMATION FOR SEQ ID NO: 146:
             (i) SEQUENCE CHARACTERISTICS:
30
                    (A) LENGTH: 99 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 146:
35
      Met Lys Ile Pro Val Leu Pro Ala Val Val Leu Leu Ser Leu Leu Val
      Leu His Ser Ala Gln Gly Ala Thr Leu Gly Gly Pro Glu Glu Glu Ser
40
      Thr Ile Glu Asn Tyr Ala Ser Arg Pro Glu Ala Phe Asn Thr Pro Phe
               35
                                   40
      Leu Asn Ile Asp Lys Leu Arg Ser Ala Phe Lys Ala Asp Glu Phe Leu
45
       Asn Trp His Ala Leu Phe Glu Ser Ile Lys Arg Lys Leu Pro Phe Leu
50
       Asn Trp Asp Ala Phe Pro Lys Leu Lys Gly Leu Arg Ser Ala Thr Pro
                                            90
       Asp Ala Gln
55
       (2) INFORMATION FOR SEQ ID NO: 147:
 60
```

```
(i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 8 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
 5
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 147:
      Met Val Trp Gly Leu Leu Leu Gly
                        5
10
      (2) INFORMATION FOR SEQ ID NO: 148:
              (i) SEQUENCE CHARACTERISTICS:
15
                     (A) LENGTH: 39 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 148:
20
      Met Leu Pro Leu Leu Ser Leu Leu Phe Leu Phe Phe Ser Thr Val Ser
                                          10
      Ser Phe Cys Gly Met Pro Leu Arg Ala His Thr Arg Ala Xaa Ala His
                  20
25
      Thr Arg Thr Phe Ala Ser Arg
               35
30
      (2) INFORMATION FOR SEQ ID NO: 149:
              (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 131 amino acids
35
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 149:
      Met Ile Cys Glu Thr Lys Ala Arg Lys Ser Ser Gly Gln Pro Gly Arg
40
      Leu Pro Pro Pro Thr Leu Ala Pro Pro Gln Pro Pro Leu Pro Glu Thr
                   20
45
      Ile Glu Arg Pro Val Gly Thr Gly Ala Met Val Ala Arg Ser Ser Asp
                                   40
      Leu Pro Tyr Leu Ile Val Gly Val Val Leu Gly Ser Ile Val Leu Ile
                               55
50
      Ile Val Thr Phe Ile Pro Phe Cys Leu Trp Arg Ala Trp Ser Lys Gln
      Lys His Thr Thr Asp Leu Gly Phe Pro Arg Ser Ala Leu Pro Pro Ser
55
      Cys Pro Tyr Thr Met Val Pro Leu Gly Gly Leu Pro Gly His Gln Ala
                                      105
60
      Val Asp Ser Pro Thr Ser Val Ala Ser Val Asp Gly Pro Val Leu Met
```

287

125 120 115 Gly Ser Thr 130 5 (2) INFORMATION FOR SEQ ID NO: 150: (i) SEQUENCE CHARACTERISTICS: 10 (A) LENGTH: 32 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 150: 15 Met Gly Ala Pro Ser Leu Thr Met Leu Leu Leu Leu Lys Val Gln Pro Arg Arg Thr Gln Ala Phe Asp Ala His Trp Val Gly Leu Pro Leu Leu 20 25 25 (2) INFORMATION FOR SEQ ID NO: 151: (i) SEQUENCE CHARACTERISTICS: 30 (A) LENGTH: 14 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 151: Met Cys Leu Ile Phe Leu Leu Leu Leu Leu Ser Phe Ser 35 5 (2) INFORMATION FOR SEQ ID NO: 152: 40 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 8 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear 45 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 152: His Pro His Gln Asp Ser Gln Pro 5 1 50 (2) INFORMATION FOR SEQ ID NO: 153: (i) SEQUENCE CHARACTERISTICS: 55 (A) LENGTH: 68 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 153: 60

	1		1111	ser	5		reu	Arg	Leu	10		vai	. Val	Ser	Val 15	Vai
5	Ile	Tyr	Leu	Ala 20		His	Pro	Leu	Leu 25	Ser	Phe	Ser	Leu	Glu 30		Pro
	Leu	Leu	Val 35	Pro	Trp	Arg	Asp	Cys 40		Gln	Asn	Ile	Trp		Ser	Gly
10	Ser	Va1 50	Trp	Tyr	Lys	Arg	Trp 55	Thr	Leu	Pro	His	Met 60	Glu	Val	Cys	Cys
15	Gln 65	Asp	Leu	His												
20	(2)				FOR											
20			(i)	(ENCE A) L B) T	ENGT	H: 2	6 am	ino		ls					
25			(xi)		D) T UENC					EQ I	D NO	: 15	4:			
	Met 1	Leu	Lys	Ile	Phe 5	Lys	Glu	Trp	Glu	Asn 10	Leu	Asn	Leu	Ile	Leu 15	Thr
30	Ser	Ile	Arg	Ile 20	Leu	Glu	Arg	Gln	Asn 25	Met						
35	(2)	INFO	ORMAT	NOIT	FOR	SEQ	ID I	NO: í	155:							
			(i) :	(ENCE A) L B) T	ENGT	H: 1	95 a	mino		ds					
40			(xi)		D) T UENCI					EQ I	D NO	: 15	5:			
	Met 1	Asp	Cys	Glu	Val 5	Asn	Asn	Gly	Ser	Ser 10	Leu	Arg	Asp	Glu	Cys 15	Ile
45	Thr	Asn	Leu	Leu 20	Val	Phe	Gly	Phe	Leu 25	Gln	Ser	Cys	Ser	Asp 30	Asn	Ser
50	Phe	Arg	Arg 35	Glu	Leu	Asp	Ala	Leu 40	Gly	His	Glu	Leu	Pro 45	Val	Leu	Ala
30	Pro	Gln 50	Trp	Glu	Gly	Tyr	Asp 55	Glu	Leu	Gln	Thr	Asp 60	Gly	Asn	Arg	Ser
55	Ser 65	His	Ser	Arg	Leu	Gly 70	Arg	Ile	Glu	Ala	Asp 75	Ser	Glu	Ser	Gln	Glu 80
	Asp	Ile	Ile	Arg	Asn 85	Ile	Ala	Arg	His	Leu 90	Ala	Gln	Val	Gly	Asp 95	Ser
60	Met	Asp	Arg	Ser	Ile	Pro	Pro	Gly	Leu	Val	Asn	Glv	Leu	Ala	Leu	Gln

				100					105					110		
_	Leu	Arg	Asn 115	Thr	Ser	Arg	Ser	Glu 120	Glu	Asp	Arg	Asn	Arg 125	Asp	Leu	Ala
5	Thr	Ala 130	Leu	Glu	Gln	Leu	Leu 135	Gln	Ala	Tyr	Pro	Arg 140	Asp	Met	Glu	Lys
10	Glu 145	Lys	Thr	Met	Leu	Val 150	Leu	Ala	Leu	Leu	Leu 155	Ala	Lys	Lys	Val	Ala 160
	Ser	His	Thr	Pro	Ser 165	Leu	Leu	Arg	Asp	Val 170	Phe	His	Thr	Thr	Val 175	Asn
15	Phe	Ile	Asn	Gln 180	Asn	Leu	Arg	Thr	Tyr 185	Val	Arg	Ser	Leu	Ala 190	Arg	Asn
20	Gly	Met	Asp 195													
25	(2)	INF	(i)	SEQU	ENCE (A) I (B) I	CHA LENG LYPE LOPOI	RACTH: :	91 ar ino a : lin	STICS mino acid near	acio						
30	Met 1					ı Val			DN: S		. Gly			Thr	Leu 15	Ala
35	Cys	: Sei	: Phe	e Leu 20		, Pro	Ly:	s Alá	a Arg		Ser	. Lys	s Arg	Ser 30		Arg
	Asr	туі	Thi		Sei	Thi	: Se:	r Pro		y Gly	y Pro	Arg	g Ala 49		Arg	g Gly
40	Gly	/ Ala 5		o Arç	g Lei	ı Sei	Se:		n Gli	n Ası	n Sei	r Sei) Lys	Gly	/ Val
45	A1a		l Ala	a Lys	s Ala	a Se:		r Ar	g Pr	o Va	l Le: 7!		s Phe	e Lei	ı Pro	Gly 80
	Pr	o Tr	p Se	r Se:	r Xaa 8		o Xa	a Al	a Ph	e Le		e				
50	(2) IN	FORM	OITA	n FO	R SE	Q II	NO:	157	:						
55					(A) (B) (D)	LENG TYPI TOPG	STH: E: au OLOG	31 a mino Y: 1:	STIC amino acio inea ION:	aci i		JO: 1	L57:			
60	Me	et Gl	y Th	ır L∈	eu Se	er Al	a G	Lu Cy	,s Se		.y Pr .0	o Al	a Th	ır Le		y Leu .5

	Суѕ	Leu	Val	Val 20	Pro	Trp	Asn	Ser	Ser 25	Gly	Leu	Ser	Gln	Pro 30	Pro	
5																
	(2)	INF	ORMA!	rion	FOR	SEQ	ID I	NO: 1	158:							
10			(i) (xi)	(A) L B) T D) T	ENGT YPE : OPOL	H: 9 ami OGY:	ERIS' 1 am no a lin PTIO	ino cid ear	acid		: 15	8:			
15	Met 1	Lys	Phe	Leu	Ala 5	Val	Leu	Val	Leu	Leu 10	Gly	Val	Ser	Ile	Phe 15	Leu
20	Val	Ser	Ala	Gln 20	Asn	Pro	Thr	Thr	Ala 25	Ala	Pro	Ala	Asp	Thr 30	Tyr	Pro
	Ala	Thr	Gly 35	Pro	Ala	Asp	Asp	Glu 40	Ala	Pro	Asp	Ala	Glu 45	Thr	Thr	Ala
25	Ala	Ala 50	Thr	Thr	Ala	Thr	Thr 55	Ala	Ala	Pro	Thr	Thr 60	Ala	Thr	Thr	Ala
	Ala 65	Ser	Thr	Thr	Ala	Arg 70	Lys	Asp	Ile	Pro	Val 75	Leu	Pro	Lys	Trp	Val 80
30	Gly	Asp	Leu	Pro	Asn 85	Gly	Arg	Val	Cys	Pro 90	Xaa					
35	(2)	INFO	ORMAT	NOI	FOR	SEQ	ID I	10: í	L59:							
40			(i) : (xi)	(A) L B) T D) T	ENGT YPE : OPOL	H: 8 ami OGY:	9 am no a lin	ino cid ear	acid		: 15	9:			
45	Met 1	Ile	Ile	Ser	Leu 5	Phe	Ile	Tyr	Ile	Phe 10	Leu	Thr	Cys	Ser	Asn 15	Thr
	Ser	Pro	Ser	Tyr 20	Gln	Gly	Thr	Gln	Leu 25	Gly	Leu	Gly	Leu	Pro 30	Ser	Ala
50	Gln	Trp	Trp 35	Pro	Leu	Thr	Gly	Arg 40	Arg	Met	Gln	Cys	Cys 45	Arg	Leu	Phe
	Cys	Phe 50	Leu	Leu	Gln	Asn	Cys 55	Leu	Phe	Pro	Phe	Pro 60	Leu	His	Leu	Ile
55	Gln 65	His	Asp	Pro	Cys	Glu 70	Leu	Val	Leu	Thr	Ile 75	Ser	Trp	Asp	Trp	Ala 80
60	Glu	Ala	Gly	Ala	Ser 85	Leu	Tyr	Ser	Pro							

	(2)	INFC	RMAT	ION	FOR	SEQ	ID N	0: 1	60:							
5			(i) S	- (<i>I</i> (E	A) LE 7T (E OT (C	ENGTH (PE: (POL(i: 1° amir XGY:	74 am no ac line	mino cid ear	acio						
10	Met		(xi) Ser											Val		Ser
	1	**- 7	Phe	C1.	5	λan	1727	T.QU	Ara	10 Tle	I.eu	Len	Pro	Ser	15 Phe	Ser
15	Pne	vai	Pne	20	Cys	ASII	vai	Беа	25	110	Lea	Dea		30		
	Ser	Phe	Met 35	Ser	Arg	Val	Leu	Gln 40	Lys	Asp	Ala	Glu	Gln 45	Glu	Ser	Gln
20	Met	Arg 50	Ala	Glu	Ile	Gln	Asp 55	Met	Lys	Gln	Glu	Leu 60	Ser	Thr	Val	Asn
25	Met 65	Met	Asp	Glu	Phe	Ala 70	Arg	Tyr	Ala	Arg	Leu 75	Glu	Arg	Lys	Ile	Asn 80
25	Lys	Met	Thr	Asp	Lys 85	Leu	Lys	Thr	His	Val 90		Ala	Arg	Thr	Ala 95	Gln
30	Leu	Ala	Lys	Ile 100	Lys	Trp	Val	Ile	Ser 105	Val	Ala	Phe	Tyr	Val 110	Leu	Gln
	Ala	Ala	Leu 115	Met	Ile	Ser	Leu	Ile 120	Trp	Lys	Tyr	Tyr	Ser 125	Val	Pro	Val
35	Ala	Val 130	. Val	Pro	Ser	Lys	Trp		Thr	Pro	Leu	Asp		Leu	Val	Ala
40	Phe		Thr	Arg	Val	Ala 150		Gly	Val	Gly	/ Ile 155		Cys	Trp	Ile	Leu 160
40	Val	. Cys	s Asn	Lys	Val 165		Ala	ı Ile	· Val	. Leu 170		Pro	Phe	Ser	•	
45	(2)	IN	FORMA	ATION	1 FOF	R SEÇ	Q ID	NO:	161:	:						
50					(A) (B) (D)	L EN G TYPE TOPO	TH: : am LOGY	TERI: 45 a ino : li IPTI	mino acid near	aci		O: 1	61:			
55		t Gl; 1	у Гу:	s Lei		e Ası 5	n Il	e Va	l Il	e Ar 1		s Pro	o Lei	ı Leı	u Lei 1	
	Le	u Va	l Gl	n Cy 2		u As	n Cy	s Cy	s Ar 2		s As	n Me	t Le	а Ту: 3		n Ile
60	Ph	e Le	u As	n Il	e Hi	s As	n Il	e Hi	s Ly	s Ph	e Se	r As	n Hi	s		

292

	35	40	45
5	(2) INFORMATION FOR S	EQ ID NO: 162:	
10	(A) LEN (B) TYP (D) TOP	HARACTERISTICS: GTH: 23 amino acid E: amino acid OLOGY: linear DESCRIPTION: SEQ	
15	Met Val Ala Ser Thr Le 1 5 Thr Thr Ala Ala Thr As 20	1	eu Phe Gly Val Ala Phe Al O 15
20	(2) INFORMATION FOR SE	Q ID NO: 163:	
25	(B) TYP (D) TOP	HARACTERISTICS: GTH: 70 amino aci E: amino acid DLOGY: linear DESCRIPTION: SEQ	
30	Met Leu Met Ala Pro Va	l Val Cys Leu Se 1	r Phe Ser Pro Cys Pro Al. 0 15
	Asp Thr Ser Leu Thr Gl	y Asp Gly Leu Ly: 25	s Ala Gly Leu Glu Arg Gl 30
35	Xaa Ala Leu Val Thr Le 35	u Phe Asp Ser Va. 40	l Thr His Phe Leu Ala Hi 45
40	50 Lys Gln Thr Ala Pro Hi	55	u Ala Phe Leu Arg Ser Gl 60
45		0	
	(2) INFORMATION FOR SE (i) SEQUENCE CH		
50	(A) LENG (B) TYPE (D) TOPG	TH: 323 amino ac : amino acid DLOGY: linear ESCRIPTION: SEQ	
55	Met Leu Leu Leu Leu Le 1 5	u Leu Leu Gly Ser 10	c Gly Gln Gly Pro Gln Glr
	Val Gly Ala Gly Gln Th 20	r Phe Glu Tyr Leu 25	ı Lys Arg Glu His Ser Leu 30

Ser Lys Pro Tyr Gln Gly Val Gly Thr Gly Ser Ser Ser Leu Trp Asn

			35					40					45			
5	Leu	Met 50	Gly	Asn	Ala	Met	Val 55	Met	Thr	Gln	Tyr	Ile 60	Arg	Leu	Thr	Pro
5	Asp 65	Met	Gln	Ser	Lys	Gln 70	Gly	Ala	Leu	Trp	Asn 75	Arg	Val	Pro	Cys	Phe 80
10	Leu	Arg	Asp	Trp	Glu 85	Leu	Gln	Val	His	Phe 90	Lys	Ile	His	Gly	Gln 95	Gly
	Lys	Lys	Asn	Leu 100	His	Gly	Asp	Gly	Leu 105	Ala	Ile	Trp	Tyr	Thr 110	Arg	Asn
15	Arg	Met	Gln 115	Pro	Gly	Pro	Val	Phe 120	Gly	Asn	Met	Asp	Lys 125	Phe	Val	Gly
20	Leu	Gly 130	Val	Phe	Val	Asp	Thr 135	Tyr	Pro	Asn	Glu	Glu 140	Lys	Gln	Gln	Glu
20	Arg 145	Val	Phe	Pro	Tyr	Ile 150	Ser	Ala	Met	Val	Asn 155	Asn	Gly	Ser	Leu	Ser 160
25	Tyr	Asp	His	Glu	Arg 165	Asp	Gly	Arg	Pro	Thr 170	Glu	Leu	Gly	Gly	Cys 175	Thr
	Ala	Ile	· Val	Arg 180		Leu	His	Tyr	Asp 185	Thr	Phe	Leu	Val	Ile 190	Arg	Tyr
30	Val	Lys	Arg 195		Leu	Thr	Ile	Met 200	Met	Asp	Ile	Asp	Gly 205	Lys	His	Glu
35	Trp	Arc 210		Cys	Ile	Glu	Val 215		Gly	Val	. Arg	Leu 220		Arg	Gly	Tyr
	Тут 225		e Gly	Thr	Ser	Ser 230		Thr	Gly	Asp	235		Asp	Asn	His	Asp 240
40	Va]	Ile	e Ser	Leu	Lys 245		Ph∈	e Glu	. Leu	Th: 250		Glu	Arg	Thr	Pro 255	Glu
	Glu	ı Glu	ı Lys	260		arg	, Ast	Val	. Phe 265		ı Pro	Ser	Val	. Asp 270		Met
45	Lys	s Le	a Pro 275		ı Met	Thr	Alá	280		ı Pro	o Pro	Leu	Ser 285		Leu	ı Ala
50	Lei	1 Ph		ı Ile	e Val	l Phe	Phe 295		Le	ı Va	l Phe	300		L Ph∈	e Ala	a Ile
30	Va 30		e Gl	y Il	e Ile	e Lei 310		r Ası	ı Ly:	s Tr	p Gli 31		ı Glı	n Sei	c Arg	320
55	Ar	g Ph	е Ту	r												
	/ 2	\ T*	TEO PM	'אשדר	אז ביר	D CE	O TD	NO.	165							

	(1) SEQUENCE CHARACTERISTICS: (A) LENGTH: 321 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear															
5			(xi)					: lir		SEQ I	D NC	: 16	55:			
	Met 1	Pro	Ser	Glu	Tyr 5		Tyr	Val	Lys	Leu 10		Ser	Asp	Cys	Ser 15	Arg
10	Pro	Ser	Leu	Gln 20		Tyr	Thr	Arg	Ala 25		Ser	Lys	Met	Arg 30		Pro
15			35					40					45			Gly
	Val	Trp 50	Ile	Leu	Tyr	Ile	Leu 55	Lys	Leu	Asn	туг	Thr 60	Thr	Glu	Glu	Cys
20	Asp 65	Met	Lys	Lys	Met	His 70	Tyr	Val	Asp	Pro	Asp 75	His	Val	Lys	Arg	Ala 80
	Gln	Lys	Tyr	Ala	Gln 85	Gln	Val	Leu	Gln	Lys 90	Glu	Cys	Arg	Pro	Lys 95	Phe
25	Ala	Lys	Thr	Ser 100	Met	Ala	Leu	Leu	Phe 105	Glu	His	Arg	Tyr	Ser 110	Val	Asp
30	Leu	Leu	Pro 115	Phe	Val	Gln	Lys	Xaa 120	Pro	Lys	Asp	Ser	Glu 125	Ala	Glu	Ser
	Lys	Tyr 130	Asp	Pro	Pro	Phe	Gly 135	Phe	Arg	Lys	Phe	Ser 140	Ser	Lys	Val	Gln
35	Thr 145	Leu	Leu	Glu	Leu	Leu 150	Pro	Glu	His	Asp	Leu 155	Pro	Glu	His	Leu	Lys 160
	Ala	Lys	Thr	Суѕ	Arg 165	Arg	Cys	Val	Val	Ile 170	Gly	Ser	Gly	Gly	Ile 175	Leu
40	His	Gly	Leu	Glu 180	Leu	Gly	His	Thr	Leu 185	Asn	Gln	Phe	Asp	Val 190	Val	Ile
45	Arg	Leu	Asn 195	Ser	Ala	Pro	Val	Glu 200	Gly	Tyr	Ser	Glu	His 205	Val	Gly	Asn
	Lys	Thr 210	Thr	Ile	Arg	Met	Thr 215	Tyr	Pro	Glu	Gly	Ala 220	Pro	Leu	Ser	Asp
50	Leu 225	Glu	Tyr	Tyr	Ser	Asn 230	Asp	Leu	Phe	Val	Ala 235	Val	Leu	Phe	Lys	Ser 240
	Val	Asp	Phe	Asn	Trp 245	Leu	Gln	Ala	Met	Val 250	Lys	Lys	Glu	Thr	Leu 255	Pro
55	Phe	Trp	Val	Arg 260	Leu	Phe	Phe	Trp	Lys 265	Gln	Val	Ala	Glu	Lys 270	Ile	Pro
50	Leu	Gln	Pro 275	Lys	His	Phe	Arg	Ile 280	Leu	Asn	Pro	Val	Ile 285	Ile	Lys	Glu

```
Thr Ala Phe Xaa His Pro Ser Val Leu Arg Ala Ser Val Lys Val Leu
                             295
     Gly Ala Glu Ile Arg Thr Ser Pro Gln Ser Val Ser Leu Pro Leu Ser
5
                         310
                                             315
     Xaa
10
     (2) INFORMATION FOR SEQ ID NO: 166:
             (i) SEQUENCE CHARACTERISTICS:
15
                    (A) LENGTH: 31 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 166:
      Met Thr Leu Asp Val Gln Thr Val Val Val Phe Ala Val Ile Val Val
20
      Leu Leu Leu Val Asn Val Ile Leu Met Phe Phe Leu Gly Thr Arg
                                      25
25
      (2) INFORMATION FOR SEQ ID NO: 167:
             (i) SEQUENCE CHARACTERISTICS:
30
                    (A) LENGTH: 72 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 167:
35
      Met Leu Pro Leu Leu Phe Cys Ala Phe Cys Leu His Lys Leu Gly Pro
                                           10
      Leu Leu Phe Leu Tyr Asp Val Leu Met Xaa His Glu Ala Val Met Arg
40
                                       25
      Thr His Gln Ile Gln Leu Pro Asp Pro Glu Phe Pro Ser Gln Gln Asn
                                   40
      Gln Val Leu Asn Lys Thr Leu Phe Asn Lys Leu Lys Lys Lys Lys
45
      Lys Lys Lys Xaa Xaa Xaa Lys Lys
       65
 50
       (2) INFORMATION FOR SEQ ID NO: 168:
 55
              (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 282 amino acids
                      (B) TYPE: amino acid
                      (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 168:
 60
```

	Met 1	Ala	Ser	Arg	Gly 5		Arg	Pro	Glu	His 10	Gly	Gly	Pro	Pro	Glu 15	Le
5	Phe	Tyr	Asp	Glu 20	Thr	Glu	Ala	Arg	Lys 25	Tyr	Val	Arg	Asn	Ser 30	Arg	Ме
	Ile	Asp	Ile 35		Thr	Arg	Met	Ala 40	Gly	Arg	Ala	Leu	Glu 45	Leu	Leu	ТУ
10	Leu	Pro 50	Glu	Asn	Lys	Pro	Cys 55	Tyr	Leu	Leu	Asp	Ile 60	Gly	Cys	Gly	Th:
15	Gly 65	Leu	Ser	Gly	Ser	Tyr 70	Leu	Ser	Asp	Glu	Gly 75	His	Tyr	Trp	Val	Gl ₃
	Leu	Asp	Ile	Ser	Pro 85	Ala	Met	Leu	Asp	Glu 90	Ala	Val	Asp	Arg	Glu 95	110
20	Glu	Gly	Asp	Leu 100	Leu	Leu	Gly	Asp	Met 105	Gly	Gln	Gly	Ile	Pro 110	Phe	Lys
	Pro	Gly	Thr 115	Phe	Asp	Gly	Cys	Ile 120	Ser	Ile	Ser	Ala	Val 125	Gln	Trp	Let
25	Cys	Asn 130	Ala	Asn	Lys	Lys	Ser 135	Glu	Asn	Pro	Ala	Lys 140	Arg	Leu	Tyr	Cys
30	Phe 145	Phe	Ala	Ser	Leu	Phe 150	Ser	Val	Leu	Val	Arg 155	Gly	Ser	Arg	Ala	Va 160
	Leu	Gln	Leu	Tyr	Pro 165	Glu	Asn	Ser	Glu	Gln 170	Leu	Glu	Leu	Ile	Thr 175	Thi
35	Gln	Ala	Thr	Lys 180	Ala	Gly	Phe	Ser	Gly 185	Gly	Met	Val	Val	Asp 190	Tyr	Pro
	Asn	Ser	Ala 195	Lys	Ala	Lys	Lys	Phe 200	Tyr	Leu	Cys	Leu	Phe 205	Ser	Gly	Pro
40	Ser	Thr 210	Phe	Ile	Pro	Glu	Gly 215	Leu	Ser	Glu	Asn	Gln 220	Asp	Glu	Val	Glu
45	Pro 225	Arg	Glu	Ser	Val	Phe 230	Thr	Asn	Glu	Arg	Phe 235	Pro	Leu	Arg	Met	Ser 240
	Arg	Arg	Gly	Met	Val 245	Arg	Lys	Ser	Arg	Ala 250	Trp	Val	Leu	Glu	Lys 255	Lys
50	Glu	Arg	His	Arg 260	Arg	Gln	Gly	Arg	Glu 265	Val	Arg	Pro	Asp	Thr 270	Gln	Туг
	Thr	Gly	Arg 275	Lys	Arg	Lys	Pro	Arg 280	Phe	Xaa						
55	(2)	TNEC	ORMAT	TON.	EOP.	SEO.	TD N	īO . 1	60.							
	121		(i) s							:						
60				C	A) L	ENGTI	H: 2	3 am	ino a	acids	3					

		(xi)	(D) TO	POLO	GY:	o ac line TION	ar	Q ID	NO:	169	:			
5	Met 1	Leu	Gly 1	Lys '	Thr 1	Lys :	Phe	Gln	Ser	Tyr 10	Lys	Ser	Phe	Ser	Arg 15	Lys
10	Leu	Met	Val (Cys 20	Pro:	Ser '	Thr									
	(2)	INFC	RMAT	ION	FOR	SEQ	ID N	0: 1	70:							
15			(i) S (xi)	(<i>I</i> (I	A) LE 3) TY O) TO	NGTH PE: POLC	H: 32 amin XGY:	28 an no ac line	mino cid ear	ació		: 170):			
20	Met 1	Trp	Arg	Pro	Ser 5	Val	Leu	Leu	Leu	Leu 10	Leu	Leu	Leu	Arg	His 15	Gly
25	Ala	Gln	Gly	Lys 20	Pro	Ser	Pro	Asp	Ala 25	Gly	Pro	His	Gly	Gln 30	Gly	Arg
	Val	His	Gln 35	Ala	Ala	Pro	Leu	Ser 40	Asp	Ala	Pro	His	Asp 45	Asp	Ala	His
30	Gly	Asn 50	Phe	Gln	Tyr	Asp	His 55	Glu	Ala	Phe	Leu	Gly 60	Arg	Glu	Val	Ala
35	Lys 65	Glu	Phe	Asp	Gln	Leu 70	Thr	Pro	Glu	Glu	Ser 75	Gln	Ala	Arg	Leu	Gly 80
	Arg	Ile	Val	Asp	Arg 85	Met	Asp	Arg	Ala	Gly 90	Asp	Gly	Asp	Gly	Trp 95	Val
40	Ser	Leu	Ala	Glu 100	Leu	Arg	Ala	Trp	Ile 105		His	Thr	Gln	Gln 110	Arg	His
	Ile	Arg	Asp 115	Ser	Val	Ser	Ala	Ala 120	Trp	Asp	Thr	Tyr	Asp 125	Thr	Asp	Arg
45	Asp	Gly 130	Arg	Val	Gly	Trp	Glu 135	Glu	Leu	Arg	Asn	Ala 140	Thr	Tyr	Gly	His
50	Туг 145		Pro	Gly	Glu	Glu 150		His	Asp	Val	Glu 155		Ala	Glu	Thr	Tyr 160
	Lys	: Lys	Met	Leu	Ala 165	Arg	Asp	Glu	Arg	170	Phe	Arg	Val	Ala	Asp 175	
55	Asp	Gly	Asp	Ser 180		Ala	Thr	Arg	185		Leu	Thr	` Ala	Phe 190		His
	Pro	Glu	1 Glu 195		Pro	His	Met	200) Ile	· Val	. Ile	205		Thr	Leu
60	Glu	ı Asp) Leu	Asp	Arg	Asn	Lys	Asp	Gl;	у Тух	. Val	Glr	ı Val	Glu	Glu	Tyr

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		210					215					220				
5	11e 225	Ala	Asp	Leu	Tyr	Ser 230	Ala	Glu	Pro	Gly	Glu 235	Glu	Glu	Pro	Ala	Trp 240
,	Val	Gln	Thr	Glu	Arg 245	Gln	Gln	Phe	Arg	Asp 250	Phe	Arg	Asp	Leu	Asn 255	Lys
10	Asp	Gly	His	Leu 260	Asp	Gly	Ser	Glu	Val 265	Gly	His	Trp	Val	Leu 270	Pro	Pro
	Ala	Gln	Asp 275	Gln	Pro	Leu	Val	Glu 280	Ala	Asn	His	Leu	Leu 285	His	Glu	Ser
15	qzA	Thr 290	Asp	Lys	Asp	Gly	Arg 295	Leu	Ser	Lys	Ala	Xaa 300	Ile	Leu	Gly	Asn
20	Trp 305	Asn	Met	Phe	Val	Gly 310	Ser	Gln	Ala	Thr	Asn 315	Tyr	Gly	Glu	Asp	Leu 320
20	Thr	Arg	His	His	Asp 325	Glu	Leu	Xaa								
25	(2)	ज्यात र	ORMA!	PTON	FOR	SEO	TD N	JO · 1	71.							
	(2)		(i)	SEQUI	ENCE	CHAI	RACT	ERIST	rics							
30			, ,,	(: (1	B) T D) T	YPE: OPOL	ami: CGY:	9 am no ad line	cid ear							
			(xi)													
35	Met 1	Cys	Trp	Leu	Arg 5	Ala	Trp	Xaa	Gln	Ile 10	Xaa	Leu	Pro	Val	Phe 15	Xaa
	Ser	Xaa	Phe	Leu 20	Ile	Gln	Leu	Leu	Ile 25	Ser	Phe	Ser	Glu	Asn 30	Gly	Phe
40	Ile	His	Ser 35	Pro	Arg	Asn	Asn	Gln 40	Lys	Pro	Arg	Asp	Gly 45	Asn	Xaa	Glu
45	Glu	Cys 50	Ala	Val	Lys	Lys	Ser 55	Cys	Gln	Leu	Cys	Thr 60	Glu	Asp	Lys	Lys
13	Tyr 65	Met	Met	Asn	Arg											
50	(2)															
	(2)		(i)													
55				(1	A) LI B) T	ENGTI YPE :	H: 10	60 ar no ac	mino cid		ds					
			(xi)					line TION		EQ II	NO:	172	2:			
50	Met 1	Trp	Leu	Phe	Ile 5	Leu	Leu	Ser	Leu	Ala 10	Leu	Ile	Ser	qzA	Ala 15	Met

	Val	Met	Asp	Glu 20	Lys	Val	Lys	Arg	Ser 25	Phe	Val	Leu	Asp	Thr 30	Ala	Ser
5	Ala	Ile	Cys 35	Asn	Tyr	Asn	Ala	His 40	Tyr	Lys	Asn	His	Pro 45	Lys	Tyr	Trp
10	Cys	Arg 50	Gly	Tyr	Phe	Arg	Asp 55	Tyr	Cys	Asn	Ile	Ile 60	Ala	Phe	Ser	Pro
10	Asn 65	Ser	Thr	Asn	His	Val 70	Ala	Leu	Lys	Asp	Thr 75	Gly	Asn	Gln	Leu	Ile 80
15	Val	Thr	Met	Ser	Cys 85	Leu	Àsn	Lys	Glu	Asp 90	Thr	Gly	Trp	Tyr	Trp 95	Cys
	Gly	Ile	Gln	Arg 100	Asp	Phe	.ia	Arg	Asp 105	qzA	Met	Asp	Phe	Thr 110	Glu	Leu
20	Ile	Val	Thr 115		Asp	Lys	Gly	Thr 120		Pro	Met	Thr	Leu 125	Val	Trp	Glu
25	Arg	Leu 130		Gly	Thr	Lys	Pro 135		Ala	Ala	Arg	Leu 140		Lys	Leu	Ser
	Ala 145		Leu	Thr	Ala	Pro 150		Arg	Pro	Ph∈	Ser 155		Phe	Ala	Tyr	Xaa 160
30																
35	(2)	INF			JENCI (A) :	E CHA LENG TYPE	ARACT TH: : am	reris 123 dino	STICS amin acid	5:	ids					
40			(xi) SE(: li:		SEQ	ID N): 1 [°]	73:			
		t Ala 1	a Xaa	a His		e Lev	ı Leı	ı Val	l Alá	a Le	_	n Sei	r Val	l Pro	His 15	Cys
45	Pr	o Hi:	s Le	u Let 20		u Glı	ı Glı	u Hi:	s Ly:		u Cy	s Ly:	s Va	l Ser 30		Phe
50	Se	r Gl	y Va 3		r Lei	u Vai	l Th	r Se: 4		g Gl	n As	p Se	r Se 4		: Туі	val
	Pr		1 Gl 0	n Th	r Le	u Ph	e Il 5		s Le	u Gl	y Pr	o Tr 6		a Trī	a Ası) Leu
55		a Pr 5	о Су	s Th	r Al	a Gl 7		p Pr	o Gl	u Al		u Ar 5	g Se	r Le	u Ar	g Leu 80
	Су	s Hi	s Se	er Hi		u Al 5	a Ar	g Xa	a As		il Se 90	er Pr	o Se	r Gl	n Al	a Ala 5
60	G]	u Gl	y Xa	a Xa	a Xa	a Ar	g Gl	у Су	's Gl	n Hi	s Aı	g Gl	y Se	r Ar	g Gl	u Le

				100					105					110		
5	Thr	Phe	Leu 115	Ser	Ala	Glu	Asn	Glu 120	Ala	Gly	Ile					
10	(2)	INF	ORMA:													
10			(i) :	((A) L B) T D) T	ENGT YPE : OPOL	H: 1 ami OGY:	29 a no a lin	mino cid ear	aci						
15			(xi)	SEQ	UENC:	E DE	SCRI	PTIO	N:S	EQ I	D NO	: 17	4:			
	Met 1	Lys	Val	Gly	Ala 5	Arg	Ile	Arg	Val	Lys 10	Met	Ser	Val	Asn	Lys 15	Ala
20	His	Pro	Val	Val 20	Ser	Thr	His	Trp	Arg 25	Trp	Pro	Ala	Glu	Trp 30	Pro	Gln
	Met	Phe	Leu 35	His	Leu	Ala	Gln	Glu 40	Pro	Arg	Thr	Glu	Val 45	Lys	Ser	Arg
25	Pro	Leu 50	Gly	Leu	Ala	Gly	Phe 55	Ile	Arg	Gln	Asp	Ser 60	Lys	Thr	Arg	Lys
30	Pro 65	Leu	Glu	Gln	Glu	Thr 70	Ile	Met	Ser	Ala	Ala 75	Asp	Thr	Ala	Leu	Trp 80
50	Pro	Tyr.	Gly	His	Gly 85	Asn	Arg	Glu	His	Gln 90	Glu	Asn	Glu	Leu	Gln 95	Lys
35	Tyr	Leu	Gln	Tyr 100	Lys	Asp	Met	His	Leu 105	Leu	Asp	Ser	Gly	Gln 110	Ser	Leu
	Gly	His	Thr 115	His	Thr	Leu	Gln	Gly 120	Ser	His	Asn	Leu	Thr 125	Ala	Leu	Asn
40	Ile															
45	(2)	INF	ORMAT	rion	FOR	SEQ	ID i	1 0: 1	L75 :							
			(i) :	(A) L	ENGT	RACTI H: 3	72 a	mino		ds					
50			(xi)	(D) T	OPOL	OGY:	lin	ear	EQ I	ON C	: 17	5:			
55	Met 1	Ala	Tyr	His	Ser 5	Phe	Leu	Val	Glu	Pro 10	Ile	Ser	Cys	His	Ala 15	Trp
رر	Asn	Lys	Asp	Arg 20	Thr	Gln	Ile	Ala	Ile 25	Cys	Pro	Asn	Asn	His 30	Glu	Val
60	His	Ile	Туг 35	Glu	Lys	Ser	Gly	Ala 40	Lys	Trp	Thr	Lys	Val 45	His	Glu	Leu

	Lys	Glu 50	His	Asn	Gly	Gln	Val 55	Thr	Gly	Ile	Asp	Тгр 60	Ala	Pro	Glu	Ser
5	Asn 65	Arg	Ile	Val	Thr	Cys 70	Gly	Thr	Asp	Arg	Asn 75	Ala	Tyr	Val	Trp	Thr 80
10	Leu	Lys	Gly	Arg	Thr 85	Trp	Lys	Pro	Thr	Leu 90	Val	Ile	Leu	Arg	Ile 95	Asn
10	Arg	Ala	Ala	Arg 100	Cys	Val	Arg	Trp	Ala 105	Pro	Asn	Glu	Asn	Lys 110	Phe	Ala
15	Val	Gly	Ser 115	Gly	Ser	Arg	Val	Ile 120	Ser	Ile	Cys	Tyr	Phe 125	Glu	Gln	Glu
	Asn	Asp 130	Trp	·Trp	Val	Cys	Lys 135	His	Ile	Lys	Lys	Pro 140	Ile	Arg	Ser	Thr
20	Val 145	Leu	Ser	Leu	Asp	Trp 150	His	Pro	Asn	Asn	Val 155	Leu	Leu	Ala	Ala	Gly 160
25	Ser	Cys	Asp	Phe	Lys 165	Cys	Arg	Ile	Phe	Ser 170	Ala	Tyr	Ile	Lys	Glu 175	Val
23	Glu	Glu	Arg	Pro 180	Ala	Pro	Thr	Pro	Trp 185	Gly	Ser	Lys	Met	Pro 190	Phe	Gly
30	Glu	Leu	Met 195		Glu	Ser	Ser	Ser 200	Ser	Cys	Gly	Trp	Val 205	His	Gly	Val
	Cys	Ph∈ 210		Ala	Ser	Gly	Ser 215		Val	Ala	Trp	Val 220	Ser	His	Asp	Ser
35	Thr 225		Cys	. Leu	Ala	Asp 230		. Asp	Lys	Lys	Met 235		Val	Ala	Thr	Leu 240
40	Ala	Ser	Glu	ı Thr	Leu 245		Leu	Leu	Ala	Leu 250		Phe	Ile	Thr	Asp 255	Asn
-10	Ser	Let	ı Val	1 Ala 260		. Gly	His	asp	Cys 265		e Pro	Val	. Leu	Phe 270		Tyr
45	Asp	Ala	a Ala 275		Gly			Ser 280		e Gly	/ Gly	/ Arg	285		Val	Pro
	Lys	3 Gli 29		r Ser	Glr	n Arg	g Gly 295	_	ı Thr	r Ala	a Arg	300 300		J Ph∈	e Glr	Asn
50	Le:		p Ly:	s Lys	s Ala	310		r Glu	ı Gly	y Gl	y Thi 31		a Ala	a Gly	/ Ala	320
55	Le	u As	p Se	r Le	a His 32		s Ası	n Sei	r Vai	33		n Ile	e Sei	r Val	335	ı Ser
55	Gl	y Gl	у Lу	s Ala 34		s Cy:	s Se	r Gli	n Pho		s Th	r Th	r Gly	y Met 350		Gly
60	Gl	у Ме	t Se 35		e Tr	p As	p Va	1 Ly:		r Le	u Gl	u Se	r Ala 36		u Ly:	s Asp

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	Leu	Lys 370	Ile	Lys												
5																
	(2)	INF	OR M A	rion	FOR	SEQ	ID I	.: OV	176:	•						
10			(i) :	(A) L B) T D) T	ENGT YPE: OPOL	H: 2 ami OGY:	16 a no a lin		aci		: 17	6:			
15	Met 1	Trp	Ser	Ile	Gly 5	Ala	Gly	Ala	Leu	Gly 10	Ala	Ala	Ala	Leu	Ala 15	Leu
20	Leu	Leu	Ala	Asn 20	Thr	Asp	Val	Phe	Leu 25	Ser	Lys	Pro	Gln	Lys 30	Ala	Ala
_0	Leu	Glu	Tyr 35	Leu	Glu	Asp	Ile	Asp 40	Leu	Lys	Thr	Leu	Glu 45	Lys	Glu	Pro
25	Arg	Thr 50	Phe	Lys	Ala	Lys	Glu 55	Leu	Trp	Glu	Lys	Asn 60	Gly	Ala	Val	Ile
	Met 65	Ala	Val	Arg	Arg	Pro 70	Gly	Суѕ	Phe	Leu	Суs 75	Arg	Glu	Glu	Ala	Ala 80
30	Asp	Leu	Ser	Ser	Leu 85	Lys	Ser	Met	Leu	Asp 90	Gln	Leu	Gly	Val	Pro 95	Leu
35	Tyr	Ala	Val	Val 100	Lys	Glu	His	Ile	Arg 105	Thr	Glu	Val	Lys	Asp 110	Phe	Gln
	Pro	Туг	Phe 115	Lys	Gly	Glu	Ile	Phe 120	Leu	Asp	Glu	Lys	Lys 125	Lys	Phe	Tyr
40	Gly	Pro 130	Gln	Arg	Arg	Lys	Met 135	Met	Phe	Met	Gly	Phe 140	Ile	Arg	Leu	Gly
	Val 145	Trp	Tyr	Asn	Phe	Phe 150	Arg	Ala	Trp	Asn	Gly 155	Gly	Phe	Ser	Gly	Asn 160
45	Leu	Glu	Gly	Glu	Gly 165	Phe	Ile	Leu	Gly	Gly 170	Val	Phe	Val	Val	Gly 175	Ser
50	Gly	Lys	Gln	Gly 180	Ile	Leu	Leu	Glu	His 185	Arg	Glu	Lys	Glu	Phe 190	Gly	Asp
50	Lys	Val	Asn 195	Leu	Leu	Ser	Val	Leu 200	Glu	Ala	Ala	Lys	Met 205	Ile	Lys	Pro
55	Gln	Thr 210	Leu	Ala	Ser	Glu	Lys 215	Lys								
60	(2)	INF	ORMA'	rion	FOR	SEQ	ID I	NO: :	177 :							

	(i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 55 amino acids(B) TYPE: amino acid	
5	(D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 177:	
	Net Lys Pro Val Ser Arg Arg Thr Leu Asp Trp Ile Tyr Ser Val Leu 1 5 10 15	
10	eu Leu Ala Ile Val Leu Ile Ser Trp Gly Cys Ile Ile Tyr Ala Ser 20 25 30	
15	Met Val Ser Ala Arg Arg Gln Leu Arg Lys Lys Tyr Pro Asp Lys Ile 35 40 45	
	Phe Gly Thr Asn Glu Asn Leu 50 55	
20	(2) INFORMATION FOR SEQ ID NO: 178:	
25	 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 178: 	
30	Met Ala Ala Asn Thr Phe Val Leu Ile Met Gly Ile Pro Thr Ser Ala 1 5 10 15	L
	Asn Ala Xaa Arg Asp Leu Phe 20	
35	(2) INFORMATION FOR SEQ ID NO: 179:	
40	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 103 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 179:	
45	Met Ser Ile Cys His Arg Gly Thr Gly Ile Ala Leu Ser Ala Gly Va 1 5 10 15	1
50	Ser Leu Phe Gly Met Ser Ala Leu Leu Leu Pro Gly Asn Phe Glu Se 20 25 30	r
50	Tyr Leu Glu Leu Val Lys Ser Leu Cys Leu Gly Pro Ala Leu Ile Hi 35 40 45	s
55	Thr Ala Lys Phe Ala Leu Val Phe Pro Leu Met Tyr His Thr Trp As 50 55 60	n
	Gly Ile Arg His Leu Met Trp Asp Leu Gly Lys Gly Leu Lys Ile Pr 65 70 75	0
60	Gln Leu Tyr Gln Ser Gly Val Val Leu Val Leu Thr Val Leu Se	r

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85 90 95 Ser Met Gly Leu Ala Ala Met 100 5 (2) INFORMATION FOR SEQ ID NO: 180: 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 48 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 180: 15 Met Thr Lys Ala Ser Ser Leu Trp Pro Leu Lys Thr Thr Cys Gln Ile 10 Ser Gly Thr Val Phe Phe Phe Leu Phe Leu Phe Ser Cys Phe Leu Met 20 25 Gln Ala Gln Cys Asp Lys Phe Val Gly Trp Asp Phe Phe Phe Leu 40 25 30 (2) INFORMATION FOR SEQ ID NO: 181: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 96 amino acids (B) TYPE: amino acid 35 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 181: Met Arg Arg Ala Leu Ile Pro Pro Cys Arg Gly Gly Pro Ser Ala Ser 40 Asp Xaa Cys Cys Ser Cys Ser Pro Ser Gly Phe Ser Ala Gly Arg Gly Arg Cys Pro Val Gln Gly Cys Leu Arg Pro His Arg Val Gln Leu Leu 45 40 Arg Arg Trp Gly Pro Gly Ser Pro Ala Gly Gln Arg Leu Ser Lys Gly 50 Phe Gln Leu Leu Arg Trp Trp Gly Pro Gly Ser Pro Ala Pro Glu Pro 70 Arg Lys Gly Pro Phe Pro Pro Pro Asp Pro Pro Trp Pro Val Thr Leu 85 90 55

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	(2) INFORM	MATION I	FOR SEQ	ID NO): 18	32:							
5		(A (E (C	NCE CHAI () LENGT! () TYPE: () TOPOL (ENCE DE:	H: 95 amin OGY:	ami o ac line	no a id ar			182	2:			
10	Met Leu G 1	lu Thr	Thr Lys 5	His \	/al +	Gln	Ile 10	Ala	Cys	Met	Leu	Leu 15	Leu
	Thr Cys G	In Ile	Phe Leu	Pro S	Ser	Ser 25	Leu	Ser	Pro	Ser	Phe 30	Ile	His
15	Ser Leu T	hr Asp 35	Ser Phe	Ile	Pro 40	Leu	Lys	Lys	Leu	Tyr 45	Val	Cys	Phe
20	Val Gln S 50	Ser Thr	Leu Leu	Lys . 55	Ala	Ala	Gly	Tyr	Lys 60	Ser	Ile	Ser	Glu
20	Ala Leu G 65	Gly Phe	Asp Xaa 70		Leu	Cys	Ser	Ser 75	Ala	Arg	Phe	Val	Trp 80
25	Ile Cys H	His Thr	Tyr Ser 85	Arg	Pro	Leu	Val 90	Thr	Cys	Ala	Leu	His 95	
30	(2) INFOF	i) SEQU (ENCE CHA A) LENG B) TYPE	ARACTE TH: 2' : amin	ERIS' 7 am no a	rics ino c i d		łs					
35		xi) SEQ	D) TOPO: UENCE DI	ESCRI	PTIO	N: S							
	Met Ser '		5				10	1		Lei	ı Gly	Pro 15	Gly
40	Gly Val	Ser Met 20		ı Lys	Lys	Lys 25		ı Trp)				
45	(2) INFO	ORMATION	I FOR SE	QID!	NO:	184:							
50			DUENCE CH (A) LENC (B) TYPE (D) TOPC	TH: 1 E: ami OLOGY:	l ar ino a : lin	mino acid near	aci		0: 1	84:			
55	Met Ser 1	Gly Gl	y Leu Se 5	r Phe	. Lei	ı Lev	ı Le		1				
	(2) INFO	ORMATIO	N FOR SE	Q ID	NO:	185	:						
60		(i) SEQ	UENCE CI	HARAC	reri	STIC	S:						

```
(A) LENGTH: 65 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 185:
 5
      Met Phe Ala Asp Phe Ile Val Val Thr Ala Thr Val Gln Arg Cys Pro
      Gly Ser Pro Pro Leu Ser Glu Ile Leu Trp Lys Asp Glu Pro Phe Ala
10
                   20
                                       25
      Ile Ser Ser His Ala Gly Leu Pro Trp Leu Ser Ser Trp Pro Ala Pro
                                   40
15
      Pro Trp Thr Trp Ser Trp Ile Ser Arg Arg Arg Glu His Gly Arg Gly
                               55
      Ser
       65
20
      (2) INFORMATION FOR SEQ ID NO: 186:
25
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 22 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 186:
30
      Met Val Glu Ser Val Met Pro Val Val Val Cys Thr Leu Ser Pro Gly
                       5
                                           10
      Ile Asp Ser Ser Pro Ser
35
                   20
      (2) INFORMATION FOR SEQ ID NO: 187:
40
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 132 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
45
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 187:
      Met Asp Val Leu Phe Val Ala Ile Phe Ala Val Pro Leu Ile Leu Gly
50
      Gln Glu Tyr Glu Asp Glu Glu Arg Leu Gly Glu Asp Glu Tyr Tyr Gln
                   20
                                       25
      Val Val Tyr Tyr Tyr Thr Val Thr Pro Ser Tyr Asp Asp Phe Ser Ala
                                   40
55
      Asp Phe Thr Ile Asp Tyr Ser Ile Phe Glu Ser Glu Asp Arg Leu Asn
                               55
                                                   60
      Arg Leu Asp Lys Asp Ile Thr Glu Ala Ile Glu Thr Thr Ile Ser Leu
60
       65
                           70
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Glu Thr Ala Arg Ala Asp His Pro Lys Pro Val Thr Val Lys Pro Val
     Thr Thr Glu Pro Gln Ser Pro Asp Leu Asn Asp Ala Val Ser Ser Leu
 5
                                    105
                 100
      Arg Ser Pro Ile Pro Leu Leu Ser Cys Ala Phe Val Gln Val Gly
                                 120
10
      Met Tyr Phe Met
        130
15
      (2) INFORMATION FOR SEQ ID NO: 188:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 69 amino acids
20
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 188:
      Met Pro Cys Gln Pro Gly Gln Val Pro Ser Cys Gln Cys Thr Phe Gly
25
      Leu Leu Met Leu Pro Ser Leu Pro Ser Pro Ala Ser Gln Pro Arg
      Pro Phe Cys Ser Ser Met Glu Tyr Phe His Gly Cys Ala Ser Pro Ser
30
      Gln Ala Ile Ile Gly Gly Phe Pro Phe Ala Ser Val Ala Leu Ala Asp
                               55
35
      Ile Leu Cys Leu Gln
       65
40
       (2) INFORMATION FOR SEQ ID NO: 189:
              (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 45 amino acids
45
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 189:
      Met Ser Leu Leu Ser Pro Ala Ile Pro Ala Leu Thr Leu Ile Phe Ile
 50
       Leu Met Phe Phe Ser Phe Pro Phe Arg Ala His Thr Val Val Thr Ile
                              25
 55
       Val Ala Ser Gly Phe Leu Gly Leu Ser Pro Leu Cys Gly
                                    40
                35
       (2) INFORMATION FOR SEQ ID NO: 190:
```

```
(i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 65 amino acids
                     (B) TYPE: amino acid
 5
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 190:
      Met Ala Phe Gly Leu Gln Met Phe Ile Gln Arg Lys Phe Pro Tyr Pro
                                           10
10
      Leu Gln Trp Ser Leu Leu Val Ala Val Val Ala Gly Ser Val Val Ser
                                       25
                                                           30
      Tyr Gly Val Thr Arg Val Glu Ser Glu Lys Cys Asn Asn Leu Trp Leu
15
      Phe Leu Glu Thr Gly Gln Leu Pro Lys Asp Arg Ser Thr Asp Gln Arg
                              55
20
      Ser
       65
25
      (2) INFORMATION FOR SEQ ID NO: 191:
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 50 amino acids
                    (B) TYPE: amino acid
30
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 191:
      Met Asn Leu Leu Gly Met Ile Phe Ser Met Cys Gly Leu Met Leu Lys
35
      Leu Lys Trp Cys Ala Trp Val Ala Val Tyr Cys Ser Phe Ile Ser Phe
                                      25
      Ala Asn Ser Arg Ser Ser Glu Asp Thr Lys Gln Met Met Ser Ser Phe
40
                                   40
      Met Xaa
           50
45
      (2) INFORMATION FOR SEQ ID NO: 192:
             (i) SEQUENCE CHARACTERISTICS:
50
                    (A) LENGTH: 170 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 192:
55
      Met Leu Leu Asn Val Ala Leu Val Ala Leu Val Leu Leu Gly Ala Tyr
      Arg Leu Trp Val Arg Trp Gly Arg Arg Gly Leu Gly Ala Gly Ala Gly
                   20
                                       25
60
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	Ala	Gly	Glu 35	Glu	Ser	Pro	Ala	Thr 40	Ser	Leu	Pro	Arg	Met 45	Lys	Lys	Arg
5	Asp	Phe 50	Ser	Leu	Glu	Gln	Leu 55	Arg	Gln	Tyr	Asp	Gly 60	Ser	Arg	Asn	Pro
	Arg 65	Ile	Leu	Leu	Ala	Val 70	Asn	Gly	Lys	Val	Phe 75	Asp	Val	Thr	Lys	Gly 80
10	Ser	Lys	Phe	Tyr	Gly 85	Pro	Ala	Gly	Pro	Tyr 90	Gly	Ile	Phe	Ala	Gly 95	Arg
15	Asp	Ala	Ser	Arg 100	Gly	Leu	Ala	Thr	Phe 105	Cys	Leu	Asp	Lys	Asp 110	Ala	Leu
13	Arg	Asp	Glu 115		Asp	Asp	Leu	Ser 120	Asp	Leu	Asn	Ala	Val 125	Gln	Met	Glu
20	Ser	Val 130		Glu	Trp	Glu	Met 135		Phe	Lys	Glu	Lys 140		Asp	Tyr	Val
	Gly 145		Leu	Leu	Lys	Pro 150		Glu	Glu	Pro	Ser 155		Tyr	Thr	Asp	Glu 160
25	Glu	Asp	Thr	Lys	Asp 165		Asn	Lys	Gln	170						
30	(2)	INF	ORMA	MOITA	1 FOF	R SEÇ) ID	NO:	193:							
			(i)	SEQ	(A)	LENG'	TH:	reris 66 a ino	mino	aci	ds					
35			(xi) SE	(D)	TOPO:	LOGY	: li	near		ID N	O: 1	93:			
40		Th:	r Ty:	r Phe		r Gly 5	y Le	u Lei	u Va	l Ile		ı Ala	a Phe	⊇ Alā	Ala 15	a Trp
40	Va.	l Al	a Le	u Ala		u Gly	y Le	u Gl	y Va 2		a Vai	l Ty	r Ala	a Ala 30		a Val
45	Le	u Le		y Al	a Gl	у Су:	s Al	a Th		e Le	u Va	l Th	r Se:		ı Ala	a Met
	Th		a As O	p Le	u Il	e Gl		o Hi 5	s Th	r As	n Se	r Gl 6		u Se	r Cy:	s Thr
50	Al 6	a Pr 5	0													
55	(2	1 I (:	IFORI	1ATIC	N FC	R SE	Q II	NO:	194	: :						
			(i) SE((A)	LEN	GTH:	TERI 92 a	amin	o ac	ids					
60					(D)			Y: 1								

			(xi)	SEQ	UENC	E DE	SCRI	PTIO	N: S	EQ I	D NO	: 19	4:			
5	Met 1	Ala	Ala	Gly	Pro 5	Ser	Gly	Cys	Leu	Val 10	Pro	Ala	Phe	Gly	Leu 15	Ar
	Leu	Leu	Leu	Ala 20	Thr	Val	Leu	Gln	Ala 25	Val	Ser	Ala	Phe	Gly 30	Ala	Gl
10	Phe	Ser	Ser 35	Glu	Ala	Cys	Arg	Glu 40	Leu	Gly	Phe	Ser	Ser 45	Asn	Leu	Lei
	Cys	Ser 50	Ser	Cys	Asp	Leu	Leu 55	Gly	Gln	Phe	Asn	Leu 60	Leu	Gln	Leu	Ası
15	Pro 65	Asp	Cys	Arg	Gly	Cys 70	Cys	Gln	Glu	Glu	Ala 75	Gln	Phe	Glu	Thr	80 Lys
20	Lys	Leu	Tyr	Ala	Gly 85	Ala	Ile	Leu	Glu	Val 90	Cys	Gly				
	(2)	INFO	ORMA'	NOIT	FOR	SEQ	ID 1	1 0: [195 :							
25			(i) :	(ENCE A) L B) T D) T	ENGT YPE :	H: 1 ami	76 a no a	mino cid		ds					
20			(xi)		UENC					EQ I	D NO	: 19	5 :			
30	Met 1	Arg	Gly	Ser	His 5	Leu	Arg	Leu	Leu	Pro 10	Tyr	Leu	Val	Ala	Ala 15	Asr
35	Pro	Val	Asn	Tyr 20	Gly	Arg	Pro	Tyr	Arg 25	Leu	Ser	Cys	Val	Glu 30	Ala	Phe
	Ala	Ala	Thr 35	Phe	Cys	Ile	Val	Gly 40	Phe	Pro	Asp	Leu	Ala 45	Val	Ile	Let
40	Leu	Arg 50	Lys	Phe	Lys	Trp	Gly 55	Lys	Gly	Phe	Leu	Asp 60	Leu	Asn	Arg	Glr
45	Leu 65	Leu	Asp	Lys	Tyr	Ala 70	Ala	Cys	Gly	Ser	Pro 75	Glu	Glu	Val	Leu	Glr 80
	Ala	Glu	Gln	Glu	Phe 85	Leu	Ala	Asn	Ala	Lуs 90	Glu	Ser	Pro	Gln	Glu 95	Glu
50	Glu	Ile	Asp	Pro 100	Phe	Asp	Val	Asp	Ser 105	Gly	Arg	Glu	Phe	Gly 110	Asn	Pro
	Asn	Arg	Pro 115	Val	Ala	Ser	Thr	Arg 120	Leu	Pro	Ser	Asp	Thr 125	Asp	Asp	Ser
55	Asp	Ala 130	Ser	Glu	Asp	Pro	Gly 135	Pro	Xaa	Ala	Glu	Arg 140	Gly	Gly	Ala	Ser
60	Ser 145	Ser	Cys	Cys	Glu	Glu 150	Glu	Gln	Thr	Gln	Gly 155	Arg	Gly	Ala	Glu	Ala 160

Arg Ala Pro Ala Glu Val Trp Lys Gly Ile Lys Lys Arg Gln Arg Asp 170 5 (2) INFORMATION FOR SEQ ID NO: 196: 10 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 70 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 196: 15 Met Ser Asn Ala Cys Lys Glu Leu Ala Ile Phe Leu Thr Thr Gly Ile 5 10 1 Val Val Ser Ala Phe Gly Leu Pro Ile Val Phe Ala Arg Ala His Leu 20 Ile Glu Trp Gly Ala Cys Ala Leu Val Leu Thr Gly Asn Thr Val Ile 25 Phe Ala Thr Ile Leu Gly Phe Phe Leu Val Phe Gly Ser Asn Asp Asp 60 55 Phe Ser Trp Gln Gln Trp 30 (2) INFORMATION FOR SEQ ID NO: 197: 35 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 25 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 197: 40 Met Thr Leu Leu Ile Ile Phe Leu Pro Phe Xaa Phe Thr Thr Xaa Thr 5 10 1 45 Asn Ser Gly Gly Ser Phe Pro Val Arg 20 50 (2) INFORMATION FOR SEQ ID NO: 198: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 73 amino acids (B) TYPE: amino acid 55 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 198: Met Lys Gly Glu Leu Leu Pro Phe Leu Phe Leu Thr Val Trp Leu Trp

5 10

	Leu	Tyr	Lys	Leu 20	Xaa	Phe	Gly	Glu	Ser 25	Pro	Arg	Tyr	Pro	Asn 30	Val	Ile
5	Gly	Lys	Thr 35	Tyr	Phe	Phe	Phe	Trp 40	Thr	Asp	Gln	Ile	Ser 45	Arg	Glu	Ser
	Arg	Phe 50	Leu	Glu	Arg	Leu	Ala 55	Phe	Ile	Val	Ser	Glu 60	Asn	Суѕ	Leu	Ile
10	Phe 65	Leu	Ile	His	Ala	Ile 70	Thr	Gly	Gln							
15	(2)	INF	ORMA!	rion	FOR	SEQ	ID I	VO : 1	199:							
20			(i) (xi)	(A) L B) T D) T	ENGT YPE: OPOL	H: 2 ami OGY:	89 a no a lin	mino cid ear	aci		: 19	9:			
25	Met 1	Ser	Gly	Phe	Ser 5	Thr	Glu	Glu	Arg	Ala 10	Ala	Pro	Phe	Ser	Leu 15	Glu
 0	Tyr	Arg	Val	Phe 20	Leu	Lys	Asn	Glu	Lys 25	Gly	Gln	Tyr	Ile	Ser 30	Pro	Phe
30	His	Asp	Ile 35	Pro	Ile	Tyr	Ala	Asp 40	Lys	Asp	Val	Phe	His 45	Met	Val	Val
	Glu	Val 50	Pro	Arg	Trp	Ser	Asn 55	Ala	Lys	Met	Glu	Ile 60	Ala	Thr	Lys	Asp
35	Pro 65	Leu	Asn	Pro	Ile	Lys 70	Gln	Asp	Val	Lys	Lys 75	Gly	Lys	Leu	Arg	Tyr 80
40	Val	Ala	Asn	Leu	Phe 85	Pro	Tyr	Lys	Gly	Tyr 90	Ile	Trp	Asn	Tyr	Gly 95	Ala
	Ile	Pro	Gln	Thr 100	Trp	Glu	Asp	Pro	Gly 105	His	Asn	Asp	Lys	His 110	Thr	Gly
45	Cys	Cys	Gly 115	Asp	Asn	Asp	Pro	Ile 120	Asp	Val	Cys	Glu	Ile 125	Gly	Ser	Lys
	Val	Cys 130	Ala	Arg	Gly	Glu	Ile 135	Ile	Gly	Val	Lys	Val 140	Leu	Gly	Ile	Leu
50	Ala 145	Met	Ile	qzA	Glu	Gly 150	Glu	Thr	Asp	Trp	Lys 155	Val	Ile	Ala	Ile	Asn 160
55	Val	Asp	Asp	Pro	Asp 165	Ala	Ala	Asn	Tyr	Asn 170	Asp	Ile	Asn	Asp	Val 175	Lys
	Arg	Leu	Lys	Pro 180	Gly	Tyr	Leu	Glu	Ala 185	Thr	Val	Asp	Trp	Phe 190	Arg	Arg
60	Tyr	Lys	Val 195	Pro	Asp	Gly	Lys	Pro 200	Glu	Asn	Glu	Phe	Ala 205	Phe	Asn	Ala

	Glu	Phe 210	Lys	Asp	Lys	Ası		he A 15	la :	Ile	Asp	11		le I 20	Lys	Ser	Th	r H	is
5	Asp 225	His	Trp	Lys	Ala	Lei 23		al T	hr	Lys	Lys	Th 23	ır A	sn (Gly	Lys	G1;	y I 2	le 40
1.0	Ser	Cys	Met	Asn	Thr 245		r L	eu S	Ser	Glu	Ser 250		O P	he 1	Lys	Cys	As 25	рР 5	ro
10	Asp	Ala	Ala	Arc 260		a Il	e V	al A		Ala 265	Leu	ı Pı	:0 P	ro	Pro	Cys 270	Gl	u S	er
15	Ala	Cys	Thr 275		Pro	o Th	ır A		/al 280	Asp	Lys	т	rp F	he	His 285	His	Gl	n L	ys
	Asn																		
20																			
	(2)	INF	ORMA		N FO						S:								
25			(1)	SEV	(A) (B)	LEN TYP	GTH E:	: 62 amir	25 a 10 a	mino cid		ids	5						
			(xi) SE	QUEN	TOP ICE					SEQ	ID	NO:	20	0:				
30	Met		ı Ile	e Pr	o Gl	y S	er :	Leu	Cys	Lys		s V O	al :	Lys	Leu	Sei	c A	sn . 15	Asn
25	Ala	a Gli	n As		p Gl	у М	et	Gln	Arg	A1a 25		r A	Asn	Val	Thr	Ту: 3	r G O	ln	Ala
35	His	s Hi	s Va 3		er Ai	cg A	sn	Lys	Arg 40		y Gl	n V	/al	Val	Gl _y 45		r A	rg	Gly
40	Gl	_	e Ar O	g Gl	Ly Cy	ys T	hr	Val 55	Trp	Le	u Th	ır (Gly	Leu 60	. Ser	Gl	уА	la	Gly
	Ly 6		r Th	ır Va	al S	er M	i et 70	Ala	Leu	ı Gl	u G.	lu ʻ	Tyr 75	Leu	∖Va.	L Cy	s H	lis	Gly 80
45	Il	e Pr	o Cy	s T		hr I 85	eu	Asp	Gly	/ As	p As	sn 90	Ile	Arg	g Gli	n Gl	уI	eu 95	Asn
	Ly	s As	sn Le		ly P 00	he S	Ser	Pro	Gli	As 10		rg	Glu	Glu	ı As	n Va 11	11 A 10	Arg	Arg
50	1)	e A.	la G: 1:	lu V 15	al A	la 1	Ьys	Leu	Pho 12		a A	sp	Ala	Gly	/ Le 12		al (Cys	Ile
55	T		er Pl	he I	le S	Ser :	Pro	Туг 135		r Gl	ln A	.sp	Arg	Ası 140		n A	la .	Arg	Gln
		le H 45	is G	lu G	Sly F		Ser 150		ı Pr	o Pl	ne F	he	Glu 155		l Ph	ie V	al.	Asp	Ala 160
60	D	T	11	ic t	al (`\/E	Glu	G1r	n Ar	a A	/ az	al	Lvs	Gl	v L∈	eu T	yr	Lys	Lys

					165					170					175	
5	Ala	Arg	Ala	Gly 180	Glu	Ile	Lys	Gly	Phe 185	Thr	Gly	Ile	Asp	Ser 190	Glu	Tyr
	Glu	Lys	Pro 195	Glu	Ala	Pro	Glu	Leu 200	Val	Leu	Lys	Thr	Asp 205	Ser	Cys	Asp
10	Val	Asn 210	Asp	Cys	Val	Gln	Gln 215	Val	Val	Glu	Leu	Leu 220	Gln	Glu	Arg	Asp
	Ile 225	Val	Pro	Val	Asp	Ala 230	Ser	Tyr	Glu	Val	Lys 235	Glu	Leu	Tyr	Val	Pro 240
15	Glu	Asn	Lys	Leu	His 245	Leu	Ala	Lys	Thr	Asp 250	Ala	Glu	Thr	Leu	Pro 255	Ala
20	Leu	Lys	Ile	Asn 260	Lys	Val	Asp	Met	Gln 265	Trp	Val	Gln	Val	Leu 270	Ala	Glu
	Gly	Trp	Ala 275	Thr	Pro	Leu	Asn	Gly 280	P'ne	Met	Arg	Glu	Arg 285	Glu	Tyr	Leu
25	Gln	Суs 290	Leu	His	Phe	Asp	Cys 295	Leu	Leu	Asp	Gly	Gly 300	Val	Ile	Asn	Leu
	Ser 305	Val	Pro	Ile	Val	Leu 310	Thr	Ala	Thr	His	Glu 315	Asp	Lys	Glu	Arg	Leu 320
30	Asp	Gly	Cys	Thr	Ala 325	Phe	Ala	Leu	Met	Tyr 330	Glu	Gly	Arg	Arg	Val 335	Ala
35	Ile	Leu	Arg	Asn 340	Pro	Glu	Phe	Phe	Glu 345	His	Arg	Lys	Glu	Glu 350	Arg	Cys
	Ala	Arg	Gln 355	Trp	Gly	Thr	Thr	Cys 360	Lys	Asn	His	Pro	Туr 365	Ile	Lys	Met
40	Val	Met 370	Glu	Gln	Gly	Asp	Trp 375	Leu	Ile	Gly	Gly	Asp 380	Leu	Gln	Val	Leu
	Asp 385	Arg	Val	Tyr	Trp	Asn 390	Asp	Gly	Leu	Asp	Gln 395	Tyr	Arg	Leu	Thr	Pro 400
45	Thr	Glu	Leu	Lys	Gln 405	Lys	Phe	Lys	Asp	Met 410	Asn	Ala	Asp	Ala	Val 415	Phe
50	Ala	Phe	Gln	Leu 420	Arg	Asn	Pro	Val	His 425	Asn	Gly	His	Ala	Leu 430	Leu	Met
	Gln	Asp	Thr 435	His	Lys	Gln	Leu	Leu 440	Glu	Arg	Gly	Tyr	Arg 445	Arg	Pro	Val
55	Leu	Leu 450	Leu	His	Pro	Leu	Gly 455	Gly	Trp	Thr	Lys	Asp 460	Asp	Asp	Val	Pro
	Leu 465	Met	Trp	Arg	Met	Lys 470	Gln	His	Ala	Ala	Val 475	Leu	Glu	Glu	Gly	Val 480
60	Leu	Asn	Pro	Glu	Thr	Thr	Val	Val	Ala	Ile	Phe	Pro	Ser	Pro	Met	Met

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					485					490					495	
~	Tyr	Ala	Gly	Pro 500	Thr	Glu	Val	Gln	Trp 505	His	Cys	Arg	Ala	Arg 510	Met	Val
5	Ala	Gly	Ala 515	Asn	Phe	Tyr	Ile	Val 520	Gly	Arg	Asp	Pro	Ala 525	Gly	Met	Pro
10	His	Pro 530	Glu	Thr	Gly	Lys	Asp 535	Leu	Tyr	Glu	Pro	Ser 540	His	Gly	Ala	Lys
	Val 545	Leu	Thr	Met	Ala	Pro 550	Gly	Leu	Ile	Thr	Leu 555	Glu	Ile	Val	Pro	Phe 560
15	Arg	Val	Ala	Ala	Туr 565	Asn	Lys	Lys	Lys	Lys 570	Arg	Met	Asp	Tyr	Tyr 575	Asp
20	Ser	Glu	His	His 580	Glu	Asp	Phe	Glu	Phe 585	Ile	Ser	Gly	Thr	Arg 590	Met	Arg
20	Lys	Leu	Ala 595		Glu	Gly	Gln	Lys 600	Pro	Pro	Glu	Gly	Phe 605		Ala	Pro
25	Lys	Ala 610		Thr	Val	Leu	Thr 615		Tyr	Tyr	Lys	Ser 620		Glu	Lys	Ala
	Xaa 625															
30	(2)	TNT		Λ ΩΤΤ.	T FOR	SEC	מד ו	NO:	201:							
35	(2)	INI	(i)	SEQ	JENCE (A) I (B) '	E CHA LENG TYPE TOPO	\RAC' TH: : am LOGY	TERIS 649 ino : li	STICS amino acid	3: o ac		D: 20	01:			
40		: Se:	r Alá	a Sei		n Ası	o Lei	u Glı	ı Pro	Lys 10		Let	ı Phe	e Pro	Lys	
45				20	0				u Sei 25	5				3()	
			3	5				4					4!	5		
50	Le	u Gl 5		l Ar	g Se	r Ly		r Gl 5	y Pr	o Le	u Ly:	s Pr		a Ar	g Gl	ı Asp
		r Gl 5	u As	n Ly	s As	р Ні 7		a Gl	y Gl	u Il	e Se 7		r Le	u Pr	o Ph	e Pro 80
55	Gl	y Va	l Va	l Le		s Pr 5	o Al	a Al	a Se		g Gl	y Gl	y Pr	o Gl	y Le	
60	Ly	's As	n Gl	y G1 10		u Ly	rs Ly	⁄s Gl	u As 10		g Ly	s Il	e As	p Al 11		a Ly:

	Asn	Thr	Phe 115		Ser	Lys	Ile	120		Glu	Glu	Leu	125		Gly	Thi
5	Pro	Pro 130		. Arg	Phe	Pro	Lys 135		Pro	Ser	Lys	Leu 140		· Val	Gly	Gl _y
	Pro 145	Trp	Gly	Gln	Ser	Gln 150		Lys	Glu	Lys	Gly 155		Lys	Asn	Ser	Ala 160
10	Thr	Pro	Lys	Gln	Lys 165		Leu	Pro	Pro	Leu 170	Phe	Thr	Leu	Gly	Pro 175	
15	Pro	Pro	Lys	Pro 180	Asn	Arg	Pro	Pro	Asn 185		Asp	Leu	Thr	Lys 190	Phe	His
	Lys	Thr	Ser 195	Ser	Gly	Asn	Ser	Thr 200	Ser	Lys	Gly	Gln	Thr 205	Ser	Tyr	Ser
20	Thr	Thr 210	Ser	Leu	Pro	Pro	Pro 215	Pro	Pro	Ser	His	Pro 220	Ala	Ser	Gln	Pro
	Pro 225	Leu	Pro	Ala	Ser	His 230	Pro	Ser	Gln	Pro	Pro 235	Val	Pro	Ser	Leu	Pro 240
25		Arg			245					250					255	
.30		Asn		260					265					270		
		Glu	275					280					285			
35		Glu 290					295					300				
4.0	305	Glu				310					315					320
40		Lys			325					330					335	
45		Ala		340					345					350		_
		Gly	355					360					365			
50		Trp 370					375					380				
	Thr 385	Ala	Val	Glu	Ile	Asp 390	Tyr	Asp	Ser	Leu	Lys 395	Leu	Lys	Lys	Asp	Ser 400
55	Leu	Gly	Ala	Pro	Ser 405	Arg	Pro	Ile	Glu	Asp 410	Asp	Gln	Glu	Val	Tyr 415	Asp
60	Asp	Val	Ala	Glu 420	Gln	Asp	Asp	Ile	Ser 425	Ser	His	Ser	Gln	Ser 430	Gly	Ser

	Gly	Gly	Ile 435	Phe	Pro	Pro	Pro	Pro 440	Asp	Asp	Asp	Ile	Tyr 445	Asp	Gly	Ile
5	Glu	Glu 450	Glu	Asp	Ala	Asp	Asp 455	Gly	Ser	Thr	Leu	Gln 460	Val	Gln	Glu	Lys
	Ser 465	Asn	Thr	Trp	Ser	Trp 470	Gly	Ile	Leu	Lys	Met 475	Leu	Lys	Gly	Lys	Asp 480
10	Asp	Arg	Lys	Lys	Ser 485	Ile	Arg	Glu	Lys	Pro 490	Lys	Val	Ser	Asp	Ser 495	Asp
15	Asn	Asn	Glu	Gly 500	Ser	Ser	Phe	Pro	Ala 505	Pro	Pro	Lys	Gln	Leu 510	Asp	Met
13	Gly	Asp	Glu 515	Val	Tyr	Asp	Asp	Val 520	Asp	Thr	Ser	Asp	Phe 525	Pro	Val	Ser
20	Ser	Ala 530	Glu	Met	Ser	Gln	Gly 535	Thr	Asn	Val	Gly	Lys 540	Ala	Lys	Thr	Glu
	Glu 545	Lys	Asp	Leu	Lys	Lys 550	Leu	Lys	Lys	Gln	Xaa 555	Lys	Xaa	Xaa	Lys	Asp 560
25	Phe	Arg	Lys	Lys	Phe 565	Lys	Tyr	Asp	Gly	Glu 570	Ile	Arg	Val	Leu	Tyr 575	Ser
30	Thr	Lys	Val	Thr 580	Thr	Ser	Ile	Thr	Ser 585	Lys	Lys	Trp	Gly	Thr 590	Arg	Asp
50	Leu	Gln	Val 595	Lys	Pro	Gly	Glu	Ser 600	Leu	Glu	Val	Ile	Gln 605	Thr	Thr	Asp
35	Asp	Thr 610	-	Val	Leu	Cys	Arg 615		Glu	Glu	Gly	Lys 620		Gly	Tyr	Val
	Leu 625		Ser	Tyr	Leu	Ala 630		Asn	Asp	Gly	Glu 635		Tyr	Asp	Asp	Ile 640
40	Ala	Asp	Gly	Cys	Ile 645		Asp	Asn	Asp							
45	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	202:							
50			• 1	_	(A) 1 (B) ' (D) '	LENG' I'YPE I'OPO!	TH: ! : am: LOGY	TERIS 55 ar ino a : lir	mino acid near	aci		- 0				
					_			IPTIC						_		
55	Met 1		a Trp	Pro	Ser		g Ser	. Lys	Met	: Phe 10		. Le	ı Let	ı Pro	0 Va. 15	L Leu
55	Cys	5 Туі	c Lev	1 Trp 20		. Le	ı Trţ	Lev	Pro 25		n Phe	e Sei	r Trj	30 30		n Glu
60	Le	ı Lys	s Ala	a Vai	l Lei	ı Arg	g Asp	Asp A		/ Le	u Il	e Se:	r Ala		l Ala	a Trp

	Asn	Ala 50	Glu	Phe	Gln	Thr	Суs 55									
5																
	(2)	INF	ORMA'	rion	FOR	SEQ	ID I	NO: 3	203:							
10			(i) (xi)	(A) L B) T D) T	ENGT YPE : OPOL	H: 2 ami OGY:	67 a no a lin	mino cid ear	aci		: 20	3:			
15	Met 1	Val	Lys	Val	Thr 5	Phe	Asn	Ser	Ala	Leu 10	Ala	Gln	Lys	Glu	Ala 15	Lys
20	Lys	Asp	Glu	Pro 20	Lys	Ser	Gly	Glu	Glu 25	Ala	Leu	Ile	Ile	Pro 30	Pro	Asp
20	Ala	Val	Ala 35	Val	Asp	Cys	Lys	Asp 40	Pro	Asp	Asp	Val	Val 45	Pro	Val	Gly
25	Gln	Arg 50	Arg	Ala	Trp	Cys	Trp 55	Cys	Met	Cys	Phe	Gly 60	Leu	Ala	Phe	Met
	Leu 65	Ala	Gly	Val	Ile	Leu 70	Gly	Gly	Ala	Tyr	Leu 75	Tyr	Lys	Tyr	Phe	Ala 80
30	Leu	Gln	Pro	Asp	Asp 85	Val	Tyr	Tyr	Cys	Gly 90	Ile	Lys	Tyr	Ile	Lys 95	Asp
35	Asp	Val	Ile	Leu 100	Asn	Glu	Pro	Ser	Ala 105	Asp	Ala	Pro	Ala	Ala 110	Leu	Tyr
	Gln	Thr	Ile 115	Glu	Glu	Asn	Ile	Lys 120	Ile	Phe	Glu	Glu	Glu 125	Glu	Val	Glu
40	Phe	Ile 130	Ser	Val	Pro	Val	Pro 135	Glu	Phe	Ala	Asp	Ser 140	Asp	Pro	Ala	Asn
	Ile 145	Val	His	Asp	Phe	Asn 150	Lys	Lys	Leu	Thr	Ala 155	Tyr	Leu	Asp	Leu	Asn 160
45	Leu	Asp	Lys	Cys	Tyr 165	Val	Ile	Pro	Leu	Asn 170	Thr	Ser	Ile	Val	Met 175	Pro
50	Pro	Arg	Asn	Leu 180	Leu	Glu	Leu	Leu	Ile 185	Asn	Ile	Lys	Ala	Gly 190	Thr	Tyr
50	Leu	Pro	Gln 195	Ser	Tyr	Leu	Ile	His 200	Glu	His	Met	Val	Ile 205	Thr	Asp	Arg
55	Ile	Glu 210	Asn	Ile	Asp	His	Leu 215	Gly	Phe	Phe	Ile	Tyr 220	Arg	Leu	Cys	His
	Asp 225	Lys	Glu	Thr	Tyr	Lys 230	Leu	Gln	Arg	Arg	Glu 235	Thr	Ile	Lys	Gly	Ile 240
60	Gln	Lys	Arg	Glu	Ala	Ser	Asn	Cys	Phe	Ala	Ile	Arg	His	Phe	Glu	Asn

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					245				:	250				:	255	
5	Lys	Phe		Val 260	Glu '	Thr :	Leu	Ile	Cys 265	Ser :	Xaa					
	(2)	INFC	RMAT	ION	FOR	SEQ	ID N	10: 2	04:							
10		,	(i) S	(<i>I</i>	A) LE 3) TY	NGTH	H: 33	ERIST 15 am no ac line	nino rid		ls					
15			(xi)	SEQU	JENCE	E DES	CRIE	MOITS	I: SE	Q II	NO:	204	:			
	Met 1	Asp	Leu	Arg	Gln 5	Phe	Leu	Met	Cys	Leu 10	Ser	Leu	Cys	Thr	Ala 15	Phe
20	Ala	Leu	Ser	Lys 20	Pro	Thr	Glu	Lys	Lys 25	Asp	Arg	Val	His	His 30	Glu	Pro
	Gln	Leu	Ser 35	Asp	Lys	Val	His	Asn 40	Asp	Ala	Gln	Ser	Phe 45	Asp	Tyr	Asp
25	His	Asp 50	Ala	Phe	Leu	Gly	Ala 55	Glu	Glu	Ala	Lys	Thr 60	Phe	Asp	Gln	Leu
	Thr 65	Pro	Glu	Glu	Ser	Lys 70	Glu	Arg	Leu	Gly	Lys 75	Ile	Val	Ser	Lys	Ile 80
30	Asp	Gly	Asp	Lys	Asp 85	Gly	Phe	Val	Thr	Val 90	Asp	Glu	Leu	Lys	Asp 95	Trp
35	Ile	Lys	Phe	Ala 100	Gln	Lys	Arg	Trp	Ile 105	Tyr	Glu	Asp	Val	Glu 110	Arg	Gln
	Trp	Lys	Gly 115	His	Asp	Leu	Asn	Glu 120	Asp	Gly	Leu	Val	Ser 125	Trp	Glu	Glu
40	Tyr	Lys 130		Ala	Thr	Tyr	Gly 135	Tyr	Val	Leu	Asp	Asp 140	Pro	Asp	Pro	Asp
45	Asp 145		Phe	Asn	Tyr	Lys 150	Gln	Met	Met	Val	Arg 155	Asp	Glu	Arg	Arg	Phe 160
43	Lys	Met	Ala	Asp	Lys 165	Asp	Gly	' Asp	Leu	Ile 170	Ala	Thr	Lys	Glu	Glu 175	Phe
50	Thr	Ala	. Phe	Leu 180		Pro	Glu	Glu	Tyr 185		Tyr	Met	Lys	Asp 190	Ile	Val
	Val	Glr	195		Met	Glu	. Asr	200		Lys	Asn	Ala	Asp 205	Gly	Phe	Ile
55	Asp	210		ı Glu	Tyr	Ile	Gl ₃ 215		Met	Туг	Ser	His 220		Gly	Asn	Thr
60	As ₁		ı Pro	Glu	ı Trp	230		5 Thr	Glu	Arg	g Glu 235		Phe	· Val	Glu	Phe 240

	Arg	Asp	Lys	Asn	Arg 245	Asp	Gly	Lys	Met	Asp 250	Lys	Glu	Glu	Thr	Lys 255	Asp
5	Trp	Ile	Leu	Pro 260	Ser	Asp	Tyr	Asp	His 265	Ala	Glu	Ala	Glu	Ala 270	Arg	His
	Leu	Val	Тут 275	Glu	Ser	Asp	Gln	Asn 280	Lys	Asp	Gly	Lys	Leu 285	Thr	Lys	Glu
10	Glu	Ile 290	Val	Asp	Lys	Tyr	Asp 295	Leu	Phe	Val	Gly	Ser 300	Gln	Ala	Thr	Asp
15	Phe 305	Gly	Glu	Ala	Leu	Val 310	Arg	His	Asp	Glu	Phe 315					
20	(2)		ORMAT	SEQUI))	ENCE	CHAI ENGT YPE :	RACT H: 2 ami:	ERIS' 07 a no a	TICS mino cid	: aci	ds					
25			(xi)	SEQ	UENC!	E DE:	SCRI	PTIO	N: S							
	Met 1	Phe	Asp	Ala	Val 5	Leu	Ile	Leu	Leu	Leu 10	Ile	Pro	Leu	Lys	Asp 15	Lys
30	Leu	Val	Asp	Pro 20	Ile	Leu	Arg	Arg	His 25	Gly	Leu	Leu	Pro	Ser 30	Ser	Leu
	Lys	Arg	Ile 35	Ala	Val	Gly	Met	Phe 40	Phe	Val	Met	Cys	Ser 45	Ala	Phe	Ala
35	Ala	Gly 50	Ile	Leu	Glu	Ser	Lys 55	Arg	Leu	Asn	Leu	Val 60	Lys	Glu	Lys	Thr
40	Ile 65	Asn	Gln	Thr	Ile	Gly 70	Asn	Val	Val	Tyr	His 75	Ala	Ala	Asp	Leu	Ser 80
	Leu	Trp	Trp	Gln	Val 85	Pro	Gln	Tyr	Leu	Leu 90	Ile	Gly	Ile	Ser	Glu 95	Ile
45	Phe	Ala	Ser	Ile 100	Ala	Gly	Leu	Glu	Phe 105	Ala	Tyr	Ser	Ala	Ala 110	Pro	Lys
	Ser	Met	Gln 115	Ser	Ala	Ile	Met	Gly 120	Leu	Phe	Phe	Phe	Phe 125	Ser	Gly	Val
50	Gly	Ser 130	Phe	Val	Gly	Ser	Gly 135	Leu	Leu	Ala	Leu	Val 140	Ser	Ile	Lys	Ala
55	Ile 145	Gly	Trp	Met	Ser	Ser 150	His	Thr	Asp	Phe	Gly 155	Asn	Ile	Asn	Gly	Cys 160
	Tyr	Leu	Asn	Tyr	Туг 165	Phe	Phe	Leu	Leu	Ala 170	Ala	Ile	Gln	Gly	Ala 175	Thr
60	Leu	Leu	Leu	Phe 180	Leu	Ile	Ile	Ser	Val 185	Lys	Tyr	Asp	His	His 190	Arg	Asp

	His	Gln	Arg 195	Ser	Arg	Ala	Asn	Gly 200	Val	Pro	Thr		Arg . 205	Arg .	Ala	
5																
	(2)	INF	ORMAT	CION	FOR	SEQ	ID N	10: 2	06:							
10				(1	A) LI B) T D) T	ENGTI YPE : OPOLA	H: 1: amin OGY:	96 ar no ao line	mino cid ear	acio		206	5 :			
15	Met 1	Arg	Ser	Arg	Ile 5	Arg	Glu	Phe	Asp	Ser 10	Ser	Thr	Leu	Asn	Glu 15	Ser
20	Val	Arg	Asn	Thr 20	Ile	Met	Arg	Asp	Leu 25	Lys	Ala	Val	Gly	Lys 30	Lys	Phe
20	Met	His	Val 35	Leu	Tyr	Pro	Arg	Lys 40	Ser	Asn	Thr	Leu	Leu 45	Arg	Asp	Trp
25	Asp	Leu 50		Gly	Pro	Leu	Ile 55		Cys	Val	Thr	Leu 60	Ala	Leu	Met	Leu
	Gln 65	Arg	J Asp	Ser	Ala	Asp 70		Glu	Lys	Asp	Gly 75	Gly	Pro	Gln	Phe	Ala 80
30	Glu	Val	L Phe	val	Ile 85		Trp	Phe	Gly	Ala 90	Val	Thr	Ile	Thr	Leu 95	Asn
35	Ser	Lys	: Lev	100		Gly	Asn	lle	Ser 105	Phe	Phe	Gln	Ser	Leu 110	Cys	Val
33	Leu	Gl	у Туг 115	c Cys	Ile	. Leu	Pro	Leu 120		Val	Ala	Met	Leu 125	Ile	Cys	Arg
40	Leu	13		ı Lev	ı Ala	Asp	Pro 135		Pro	Val	. Asn	Phe 140		. Val	Arg	Leu
	Phe 145		l Vai	l Ile	e Val	150		e Ala	a Trp	Ser	11e		. Ala	Ser	Thr	Ala 160
45	Phe	e Le	u Ala	a Asp	Sei 169		n Pro	o Pro	o Asr	170		, Ala	a Leu	ı Ala	Val 175	Tyr
50	Pro	o Va	1 Ph	e Lei 180		е Ту	r Ph	e Vai	l Ile 185		r Trp	Met	: Ile	Leu 190		Phe
30	Th	r Pr	o Gl 19	n Xaa	a											
55	(2	1I (NFORM	OITA	N FO	R SE	Q ID	NO:	207	:						
			(i)	SEQ					STIC amin		ids					
60									acid							

(D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 207:

5	Met 1	Ala	Lys	Asp	Gln 5	Ala	Val	Glu	Asn	Ile 10	Leu	Val	Ser	Pro	Val 15	Val
	Val	Ala	Ser	Ser 20	Leu	Gly	Leu	Val	Ser 25	Leu	Gly	Gly	Lys	Ala 30	Thr	Thr
10	Ala	Ser	Gln 35	Ala	Lys	Ala	Val	Leu 40	Ser	Ala	Glu	Gln	Leu 45	Arg	Asp	Glu
15	Glu	Val 50	His	Ala	Gly	Leu	Gly 55	Glu	Leu	Leu	Arg	Ser 60	Leu	Ser	Asn	Ser
	Thr 65	Ala	Arg	Asn	Val	Thr 70	Trp	Lys	Leu	Gly	Ser 75	Arg	Leu	Tyr	Gly	Pro 80
20	Ser	Ser	Val	Ser	Phe 85	Ala	Asp	Asp	Phe	Val 90	Arg	Ser	Ser	Lys	Gln 95	His
	Tyr	Asn	Cys	Glu 100	His	Ser	Lys	Ile	Asn 105	Phe	Arg	Asp	Lys	Arg 110	Ser	Ala
25	Leu	Gln	Ser 115	Ile	Asn	Glu	Trp	Ala 120	Ala	Gln	Thr	Thr	Asp 125	Gly	Lys	Leu
30	Pro	Glu 130	Val	Thr	Lys	Asp	Val 135	Glu	Arg	Thr	Asp	Gly 140	Ala	Leu	Leu	Val
	Asn 145	Ala	Met	Phe	Phe	Lys 150	Pro	His	Trp	Asp	Glu 1 55	Lys	Phe	His	His	Lys 160
35	Met	Val	Ąsp	Asn	Arg 165	Gly	Phe	Met	Val	Thr 170	Arg	Ser	Tyr	Thr	Val 175	Gly
	Val	Met	Met	Met 180	His	Arg	Thr	Gly	Leu 185	Tyr	Asn	Tyr	Tyr	Asp 190	Asp	Glu
40	Lys	Glu	Lys 195	Leu	Gln	Ile	Val	Glu 200	Met	Pro	Leu	Ala	His 205	Lys	Leu	Ser
45	Ser	Leu 210	Ile	Ile	Leu	Met	Pro 215	His	His	Val	Glu	Pro 220	Leu	Glu	Arg	Leu
	Glu 225	Lys	Leu	Leu	Thr	Lys 230	Glu	Gln	Leu	Lys	Ile 235	Trp	Met	Gly	Lys	Met 240
50	Gln	Lys	Lys	Ala	Val 245	Ala	Ile	Ser	Leu	Pro 250	Lys	Gly	Val	Val	Glu 255	Val
	Thr	His	Asp	Leu 260	Gln	Lys	His	Leu	Ala 265	Gly	Leu	Gly	Leu	Thr 270	Glu	Ala
55	Ile	Asp	Lys 275	Asn	Lys	Ala	Asp	Leu 280	Ser	Arg	Met	Ser	Gly 285	Lys	Lys	Asp
60	Leu	Туг 290	Leu	Ala	Ser	Val	Phe 295	His	Ala	Thr	Ala	Phe 300	Glu	Leu	Asp	Thr

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	Asp 305	Gly	Asn	Pro	Leu	Thr 310	Arg	Ile	Thr	Gly	Gly 315	Gly	Vá	al A	rg T	hr (31n 320
5	Val	Phe	Tyr	Ala	Asp 325	His	Pro	Phe	Ile	Ser 330	Xaa						
10	(2)	INFO		rion sequi	ENCE		RACTI	ERIS	TICS		ls						
15			(xi)	(B) T D) T	YPE: OPOL	ami OGY:	no a lin	cid ear			D: 2	08:				
	Met 1		Met	Gln	Leu 5	Phe	Gly	Phe	Leu	Ala 10		e Me	Ι	le ¤	Phe	Met 15	Cys
20	Trp	Val	Gly	Asp 20	Val	Tyr	Pro	Val	Тут 25		n Pro	v Va	l G	ly 1	Pro 30	Lys	Gln
	Tyr	Pro	Туг 35	Asn	Asn	Leu	Tyr	Leu 40		a Arg	g Gly	/ Gl	у А	sp :	Pro	Ser	Lys
25	Glu	ı Pro		ı Arg	Val	Val	His 55		Glı	ı Ile	9						
30	(2)) INE	FORM	AOITA	I FOF	. SEÇ) ID	NO:	209	:							
35					(A) : (B) : (D) :	LENG' TYPE TOPO	TH: : am LOGY	392 ino : li	amir acid near	o ac l		10: 1	209	:			
40		t As	p Al	a Lev		l Glu 5	ı Ası) As	p Il		s Il O	e Le	eu .	Asn	His	Glu 15	Lys
	Al	a Hi	s Ly	s Arg		o Thi	r Va	l Th		o Va 5	ıl S∈	er I	le	Tyr	Ser 30	Gly	Asp
45	Gl	u Se		1 Ala	a Se	r Hi	s Ph		a Le 0	eu Va	ıl Ti	ır A	la	Тут 45	Glu	Asp	lle
	Ly		rs Ar 50	g Le	u Ly	s As	p Se 5		u Ly	/s G]	lu As		er 60	Leu	Leu	Lys	. Lys
50		rg Il	.e Ar	g Ph	e Le		u Gl 0	u Ly	s Le	eu II		la A 75	rg	Phe	Glu	Glı	ı Glu 80
55	Tł	nr Se	er Se	er Va		y Ar 15	g Gl	u Gl	ln V		sn L 90	ys A	.la	Tyr	His	3 Ala	a Tyr 5
	A	rg G	lu V	al Cy 10		le As	sp Ar	g As		sn L 05	eu L	ys S	er	Lys	Let		o Ly s
60	M	et A	sn L	ys As	sp As	sn Se	er G	lu S	er L	eu L	ys V	al I	.eu	Asr	Gl	ı Gl	n Leu

			115					120					125			
5	Gln	Ser 130	Lys	Glu	Val	Glu	Leu 135	Leu	Gln	Leu	Arg	Thr 140	Glu	Val	Glu	Th
	Gln 145		Val	Met	Arg	Asn 150	Leu	Asn	Pro	Pro	Ser 155	Ser	Asn	Trp	Glu	Va:
10	Glu	Lys	Leu	Ser	Cys 165	Asp	Leu	Lys	Ile	His 170	Gly	Leu	Glu	Gln	Glu 175	Let
	Glu	Leu	Met	Arg 180	Lys	Glu	Cys	Ser	Asp 185	Leu	Lys	Ile	Glu	Leu 190	Gln	Lys
15	Ala	Lys	Gln 195	Thr	Asp	Pro	Tyr	Gln 200	Glu	Asp	Asn	Leu	Lys 205	Ser	Arg	Asp
20	Leu	Gln 210	Lys	Leu	Ser	Ile	Ser 215	Ser	Asp	Asn	Met	Gln 220	His	Ala	Tyr	Trp
	Glu 225	Leu	Lys	Arg	Glu	Met 230	Ser	Asn	Leu	His	Leu 235	Val	Thr	Gln	Val	Glr 240
25	Ala	Glu	Leu	Leu	Arg 245	Lys	Leu	Lys	Thr	Ser 250	Thr	Ala	Ile	Lys	Lys 255	Ala
	Cys	Ala	Pro	Val 260	Gly	Cys	Ser	Glu	Asp 265	Leu	Gly	Arg	Asp	Ser 270	Thr	Lys
30	Leu	His	Leu 275	Met	Asn	Phe	Thr	Ala 280	Thr	Tyr	Thr	Arg	His 285	Pro	Pro	Leu
35	Leu	Pro 290	Asn	Gly	Lys	Ala	Leu 295	Cys	His	Thr	Thr	Ser 300	Ser	Pro	Leu	Pro
	Gly 305	Asp	Val	Lys	Val	Leu 310	Ser	Glu	Lys	Ala	Ile 315	Leu	Gln	Ser	Trp	Thr 320
40	Asp	Asn	Glu	Arg	Ser 325	Ile	Pro	Asn	Asp	Gly 330	Thr	Cys	Phe	Gln	Glu 335	His
	Ser	Ser	Tyr	Gly 340	Arg	Asn	Ser	Leu	Glu 345	Asp	Asn	Ser	Trp	Val 350	Phe	Pro
45	Ser	Pro	Pro 355	Lys	Ser	Ser	Glu	Thr 360	Ala	Phe	Gly	Glu	Thr 365	Lys	Thr	Lys
50	Thr	Leu 370	Pro	Leu	Pro	Asn	Leu 375	Pro	Pro	Leu	His	Туг 380	Leu	Asp	Gln	His
	Asn 385	Gln	Asn	Cys	Leu	Туг 390	Lys	Asn								
55	(2)	INFO	RMAT	'ION	FOR	SEQ	ID N	J O: 2	210:							
		,	(i) S					ERIST								
60						ENGTI YPE :				acid	s					

```
(D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 210:
     Met His His His Thr Gln Leu Met Phe Ile Tyr Leu Phe Ile Tyr Leu
5
                                          10
     Phe Ile Leu Gly Val Phe Phe Phe Phe Xaa
                  20
10
      (2) INFORMATION FOR SEQ ID NO: 211:
             (i) SEQUENCE CHARACTERISTICS:
15
                    (A) LENGTH: 39 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 211:
     Met Asn Cys Ile Leu Leu Leu Tyr Leu Leu Ile Pro Thr Ile Ser Ile
20
      Ser Val Val Pro Tyr Val Ala Leu Asn Ile Lys Tyr Ile Lys Glu Cys
                  20
25
      Thr Glu Asn Ser Phe Tyr Xaa
30
      (2) INFORMATION FOR SEQ ID NO: 212:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 71 amino acids
35
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 212:
      Met Leu Leu His Leu Thr Ala Ala Phe Leu Gln Arg Ala Gln Phe Ser
40
      Thr Tyr Phe Pro Gly Tyr Phe Asp Gly Gln Tyr Trp Leu Trp Trp Val
                                        25
                                                           3.0
      Phe Leu Val Leu Gly Phe Leu Leu Phe Leu Arg Gly Phe Ile Asn Tyr
45
      Ala Lys Val Arg Lys Met Pro Glu Thr Phe Ser Asn Leu Pro Arg Thr
                               55
50
       Arg Val Leu Phe Ile Tyr Xaa
       65
 55
       (2) INFORMATION FOR SEQ ID NO: 213:
              (i) SEQUENCE CHARACTERISTICS:
                      (A) LENGTH: 83 amino acids
 60
                      (B) TYPE: amino acid
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(D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 213:
      Met Leu Thr Phe Phe Met Ala Phe Leu Phe Asn Trp Ile Gly Phe Phe
 5
      Leu Ser Phe Cys Leu Thr Thr Ser Ala Ala Gly Arg Tyr Gly Ala Ile
                   20
10
      Ser Gly Phe Gly Leu Ser Leu Ile Lys Trp Ile Leu Ile Val Arg Phe
                                   40
      Ser Thr Tyr Phe Pro Ala Phe Met Asn Ser Leu Ser Arg Ser Lys Arg
           50
                              55
15
      Thr Pro Ala Gly Ser Glu Ser Arg Cys Arg Thr Gln Arg Asn Asn His
                          70
      Leu Leu Xaa
20
      (2) INFORMATION FOR SEQ ID NO: 214:
25
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 81 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
30
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 214:
      Met Ser Lys Arg Ser Ala Ser Phe Ile Leu Leu Pro Leu Leu Phe Leu
              5
                                         10
35
      Lys Gly Ser Phe Ala Lys Leu Asn Ala Arg Ile Ser Asp Cys Leu Glu
                                     25
      Glu Arg Tyr Cys His Asn Leu Trp Met Val Phe Gln Gly Cys Val Ile
40
      Thr Glu Leu His Leu Ser Arg Met Ser Lys Thr Leu Ser Ser Leu Cys
      Tyr Asp Phe Val Ile Asn Val Tyr Ile Phe Phe Lys Phe Leu Asp Ile
45
      Thr
50
      (2) INFORMATION FOR SEQ ID NO: 215:
             (i) SEQUENCE CHARACTERISTICS:
55
                    (A) LENGTH: 49 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 215:
60
      Met Cys Ser Leu Phe Glu Ser Arg Phe Phe Cys Phe Val Leu Phe Ser
```

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	1				5					10					15	
5	Glu	Lys	Ile	Ile 20	Gln	Leu	Cys	Ala	Ser 25	Ile	Ala	Phe	Leu	Cys 30	Phe	Val
J	Lys	His	Val 35	Pro	Trp	Pro	Lys	Trp 40	Lys	Arg	Lys	Суѕ	Leu 45	Ile	Asn	Ala
10	Phe															
15	(2)		ORMAT													
20			(xi)	(A) L B) T D) T	ENGT YPE: OPOL	H: 2 ami OGY:	03 a no a lin	mino cid ear	aci		: 21	6:			
	Met 1		Leu											Leu	Leu 15	Leu
25	Leu	Leu	Ser	Ala 20	Ala	Val	Cys	Arg	Ala 25	Glu	Ala	Gly	Leu	Glu 30	Thr	Glu
30	Ser	Pro	Val 35	Arg	Thr	Leu	Gln	Val 40	Glu	Thr	Leu	Val	Glu 45	Pro	Pro	Glu
30	Pro	Суs 50	Ala	Glu	Pro	Ala	Ala 55	Phe	Gly	Asp	Thr	Leu 60	His	Ile	His	Tyr
35	Thr 65	Gly	Ser	Leu	Val	Asp 70	Gly	Arg	Ile	Ile	Asp 75	Thr	Ser	Leu	Thr	Arg 80
	Asp	Pro	Leu	Val	Ile 85	Glu	Leu	Gly	Gln	Lys 90	Gln	Val	Ile	Pro	Gly 95	Leu
40	Glu	Gln	Ser	Leu 100	Leu	Asp	Met	Cys	Val 105	Gly	Glu	Lys	Arg	Arg 110	Ala	Ile
45	Ile	Pro	Ser 115	His	Leu	Ala	Tyr	Gly 120	Lys	Arg	Gly	Phe	Pro 125	Pro	Ser	Val
	Pro	Ala 130	Asp	Ala	Val	Val	Gln 135	Tyr	Asp	Val	Glu	Leu 140	Ile	Ala	Leu	Ile
50	Arg 145		Asn	Tyr	Trp	Leu 150	Lys	Leu	Val	Lys	Gly 155	Ile	Leu	Pro	Leu	Val 160
	Gly	Met	Ala	Met	Val 165	Pro	Pro	Ser	Trp	Ala 170	Ser	Leu	Gly	Ile	Thr 175	Tyr
55	Thr	Glu	Arg	Pro 180		Asp	Pro	Lys	Ser 185	Pro	Lys	Arg	Ser	Ser 190	_	Lys
60	Arg	Asn	Glu 195	Thr	Arg	Ala	Lys	Arg 200	Asn	Asn	Lys					

	(2)	INFO	DRMAT	NOI	FOR	SEQ	ID N	IO: 2	17:							
5				() (1	A) L: B) T D) T	ENGT YPE : OPOL	H: 1: amin DGY:	86 ar no ao line		aci		: 21 ⁻	7:			
0	Met 1	Lys	Thr	Leu	Met 5	Thr	Ile	Cys	Pro	.0	Thr	Val	Leu	Leu	Val 15	Phe
15	Ser	Ile	Ser	Leu 20	Trp	Ile	Ile	Ala	Ala 25	Trp	Thr	Val	Arg	Val 30	Cys	Glu
	Ser	Pro	Glu 35	Ser	Pro	Ala	Gln	Pro 40	Ser	Gly	Ser	Ser	Leu 45	Pro	Ala	Trp
20	Tyr	His 50	Asp	Gln	Gln	Asp	Val 55	Thr	Ser	Asn	Phe	Leu 60	Gly	Ala	Met	Trp
25	Leu 65	Ile	Ser	Ile	Thr	Phe 70	Leu	Ser	Ile	Gly	Tyr 75	Gly	Asp	Met	Val	Pro 80
			-	-	85	_			Суѕ	90			-		95	_
30		_		100					Ala 105					110		
35			115					120	His				125			
55		130	_				135		Ala			140				
40	145					150			Leu Ser		155					160
	-				165				Ala	170	_,_	-7-			175	V
45				180					185							
50	(2)	INF	ORMA			_										
			(i)	((A) I	ENGT	H: 9	00 am			ls					
55			(xi)			OPOI E DE			ear N: S	EQ I	D NC): 21	.8:			
	Met 1	_	Phe	Leu	Ala 5		Leu	Val	Leu	Leu 10	_	Val	Ser	Ile	Phe 15	Leu
60	Va1	Ser	· Δla	Gln	Asn	Pro	Thr	Thr	Ala	Ala	Pro	Ala	Asn	Thr	ጥህም	Pro

30 20 25 Ala Thr Gly Pro Ala Asp Asp Glu Ala Pro Asp Ala Glu Thr Thr Ala 5 Ala Ala Thr Thr Ala Thr Thr Ala Ala Pro Thr Thr Ala Thr Thr Ala Ala Ser Thr Thr Ala Arg Lys Asp Ile Pro Val Leu Pro Lys Trp Val 10 70 75 Gly Asp Leu Pro Asn Gly Arg Val Cys Pro 85 15 (2) INFORMATION FOR SEQ ID NO: 219: (i) SEQUENCE CHARACTERISTICS: 20 (A) LENGTH: 139 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 219: 25 Met Ser Ser Ala Ala Ala Asp His Trp Ala Trp Leu Leu Val Leu Ser Phe Val Phe Gly Cys Asn Val Leu Arg Ile Leu Leu Pro Ser Phe Ser 30 Ser Phe Met Ser Arg Val Leu Gln Lys Asp Ala Glu Gln Glu Ser Gln 40 Met Arg Ala Glu Ile Gln Asp Met Lys Gln Glu Leu Ser Thr Val Asn 35 55 50 Met Met Asp Glu Phe Ala Arg Tyr Ala Arg Leu Glu Arg Lys Ile Asn 40 Lys Met Thr Asp Lys Leu Lys Thr His Val Lys Ala Arg Thr Ala Gln 90 Leu Ala Lys Ile Lys Trp Val Ile Ser Val Ala Phe Tyr Val Leu Gln 105 45 Ala Ala Leu Met Ile Ser Leu Ile Trp Lys Tyr Tyr Ser Val Pro Val 120 Ala Val Val Pro Ser Lys Trp Ile Thr Leu Xaa 50 130 135 (2) INFORMATION FOR SEQ ID NO: 220: 55 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 48 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear 60 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 220:

	Met 1	Ser	Ser	Ala	Ala 5	Ala	Asp	His	Trp	Ala 10	Trp	Leu	Leu	Val	Leu 15	Ser
5	Phe	Val	Phe	Gly 20	Cys	Asn	Val	Leu	Arg 25	Ile	Leu	Leu	Pro	Ser 30	Phe	Ser
10	Ser	Phe	Met 35	Ser	Arg	Val	Leu	Gln 40	Lys	Asp	Ala	Asp	Arg 45	Ser	His	Arg
15	(2)	INF	ORMA'	rion	FOR	SEQ	ID I	NO: 2	221:							
20	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 70 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 221:															
25	Met 1	Thr	Ala	Pro	Leu 5	Pro	Pro	Leu	Ser	Gly 10	Leu	Ala	Leu	Phe	Leu 15	Ile
	Val	Phe	Phe	Ser 20	Leu	Gly	Val	Phe	Cys 25	Ile	Cys	His	Ser	His 30	Trp	Tyr
30	His	Thr	Leu 35		Gln	Met	Ala	Gly 40	Thr	Glu	Pro	Lys	Ala 45	Leu	Leu	Leu
35	Ser	Pro 50		Ala	Ala	Thr	Thr 55	Phe	Val	Thr	Val	Thr 60		Glu	Val	Trp
	Lys 65		Gln	Ala	Leu	Ala 70										
40	(2)	INF						NO:								
45				_	(A) I (B) ((D) (LENGT TYPE : TOPOI	TH: { : am: LOGY	TERIS 33 ar ino a : lir IPTIC	nino acid near	ació		D: 22	22:			
50	Met 1		Cys	Ser	Val		ı Lev	ı Lev	ı Leu	lle 10		ı Gly	/ Leu	Arg	Cys 15	Ser
	Gly	√ Va]	l Arg	Pro 20	_	Leu	ı Val	. Gly	/ Glu 25		His	: Asr	n Pro	Ser 30		Leu
55	Va]	l Cys	Let 35		ı Leı	ı Lys	s Asp	Ser 40		Thr	Asr	ı Glr	1 Gly 45		Cys	Pro
60	Gly	7 Gly 50		o Trp	Sei	Glu	1 Arg 55		o Ile	e Glu	ı Ser	c Val		s Ser	Asp	Asn

	Суs 65	Glu	Ala	Thr	Leu	Gly 70	Tyr	Arg .	Asn	His	Ser 75	Leu	Pro	Ser	Asn	Tyr 80
5	Tyr	Asn	Ser													
10	(2)		ORMAT	SEQUE () ()	ENCE A) LI B) TY	CHAF ENGTI YPE :	RACTE H: 41 amir	ERIST 3 ami	ICS: ino a		3					
15			(xi)					line TION		EQ II	NO:	223	3:			
	Met 1	Leu	Thr	Arg	Ser 5	Leu	Lys	Thr	Leu	Pro 10	Ser	Ala	Cys	Thr	Ala 15	Phe
20	Leu	Leu	Leu	Phe 20	Phe	Leu	Phe	Ser	Ser 25	Gly	Asp	Pro	Glu	Leu 30	Ser	Cys
25	Ser	Cys	Thr 35	Leu	Arg	Thr	Gln	Ser 40	Ser	Trp	Ser					
30	(2)	INF		SEQUI () ()	ENCE A) L B) T D) T	CHAI ENGT YPE: OPOL	RACTI H: 1 ami: OGY:	ERIST 84 and no ao line	rics mino cid ear	acio						
35	Met 1		(xi) Arg					PTION Leu						Arg	His 15	Gly
40	Ala	Gln	Gly	Lys 20	Pro	Ser	Pro	Asp	Ala 25	Gly	Pro	His	Gly	Gln 30	Gly	Arg
	Val	. His	Gln 35	Ala	Ala	Pro	Leu	Ser 40	Asp	Ala	Pro	His	Asp 45	Asp	Ala	His
45	Gly	Asn 50	Phe	Gln	Tyr	Asp	His 55	Glu	Ala	Phe	Leu	Gly 60	Arg	Glu	Val	Ala
50	Lys 65		Phe	Asp	Gln	Leu 70		Pro	Glu	Glu	Ser 75	Gln	Ala	Arg	Leu	Gly 80
50	Arg	g Il∈	e Val	Asp	Arg 85		Asp	Arg	Ala	Gly 90	Asp	Gly	Asp	Gly	Trp 95	
55	Sei	Leu	ı Ala	Glu 100		Arg	Ala	Trp	Ile 105		His	Thr	Gln	Gln 110	Arg	His
	Ile	e Arg	g Asp 115		Val	Ser	· Ala	Ala 120		Asp	Thr	Tyr	Asp 125		Asp	Arg
60	Asp	o Gly	y Arg	Val	Gly	Trp	Glu	Glu	Leu	Arg	Asn	Xaa	Thr	туг	Gly	His

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		130					135					140				
5	Xaa 145	Xaa	Pro	Xaa	Glu	Glu 150	Phe	His	Asp	Val	Glu 155	Asp	Ala	Glu	Thr	Туг 160
5	L ys	Lys	Met	Leu	Xaa 165	Arg	Asp	Glu	Arg	Arg 170	Phe	Arg	Val	Ala	Asp 175	Gln
10	Asp	Gly	Asp	Ser 180	Met	Ala	Thr	Arg								
15	(2)	INF	ORMAS													
20	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 71 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 225:															
	Met	Trp	Leu											Asp	Ala	Met
25	1				5					10					15	
25	Val	Met	Asp	Glu 20	Lys	Val	Lys	Arg	Ser 25	Leu	Cys	Trp	Thr	Arg 30	Leu	Leu
30	Pro	Ser	Ala 35	Thr	Thr	Met	Pro	Xaa 40	Thr	Arg	Ile	Thr	Pro 45	Asn	Thr	Gly
30	Ala	Glu 50	Xaa	Ile	Ser	Val	Xaa 55	Thr	Ala	Thr	Ser	Ser 60	Pro	Ser	Pro	Leu
35	Thr 65	Ala	Pro	Ile	Met	Trp 70	Pro									
40	(2)	INF	ORMA			SEQ CHA				: .						
			(1)	(A) I	ENGT	H: 1	.0 am	nino		ls					
45			(xi)			OPOI E DE				EQ I	D NC): 22	6:			
	Met 1		Val	Phe	Val 5		Glu	Ile	Phe	Leu 10						
50																
	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	227:							
55			(i)	((A) I (B) T	CHA LENGT TYPE:	TH: 1	l38 a ino a	mino acid		ids					
			(xi)			E DE				SEQ I	D NC): 22	27 :			
60	Met	: Ala	Val	Ala	Thr	Leu	Ala	Ser	Glu	Thr	Leu	Pro	Leu	Leu	Ala	Leu

	1				5					10					15		
_	Thr	Phe	Ile	Thr 20	Asp	Asn	Ser	Leu	Val 25	Ala	Ala	Gly	His	Asp 30	Cys	Phe	
5	Pro	Val	Leu 35	Phe	Thr	Tyr	Asp	Ala 40	Ala	Ala	Gly	Met	Leu 45	Ser	Phe	Gly	
10	Gly	Arg 50	Leu	Asp	Val	Pro	Lys 55	Gln	Ser	Ser	Gln	Arg 60	Gly	Leu	Thr	Ala	
	Arg 65	Glu	Arg	Phe	Gln	Asn 70	Leu	Asp	Lys	Lys	Ala 75	Ser	Ser	Glu	Gly	Gly 80	
15	Thr	Ala	Ala	Gly	Ala 85	Gly	Leu	Asp	Ser	Leu 90	His	Lys	Asn	Ser	Val 95	Ser	
20	Gln	Ile	Ser	Val 100	Leu	Ser	Gly	Gly	Lys 105		Lys	Cys	Ser	Gln 110	Phe	e Cys	
20	Thr	Thr	Gly 115		Asp	Gly	Gly	Met 120		Ile	Trp	Asp	Val 125	Lys	Ser	Leu	
25	Glu	Ser 130		Leu	Lys	Asp	Leu 135		Ile	· Lys							
30	(2)	INF		ATION JOSE	JENCI (A) (B)	E CHA LENG TYPE	ARAC' TH: : am	reris 23 au ino	STIC: mino acid	S: aci	ds						
35) SE	QUEN	CE D	ESCR		ON:	SEQ							
		u Gly l	y Se	r Le		r Thi	r Al	a Pro	o Sei	r Se		a Lei	u Pro	o Th:	r Le 1	u Gly 5	
40	Ala	a Ar	g Ar	g Thi		g Se:	r Ly	s									
45	(2) IN	FORM	ATIO	n fo	R SE	Q ID	NO:	229	:							
50				SEÇ L) SI	(A) (B) (D)	TYPE TOPO	GTH: E: au OLOG	133 mino Y: 1:	amir acio inear	no ad I r		VO: 2	229:				
<i></i>	M∈	et Th 1	ır Ty	nr Ph	ne Se	er Gl 5	y L∈	eu Le	eu Va		Le Le	eu Al	la Ph	ne Al	a A	la Trp 15	-
55	Vê	al Al	la Le		La G1 20	lu Gl	ly Le	eu Gl		al Al 25	la Va	al Ty	/r Al		la A 30	la Va	
60	Le	eu Le		ly A: 35	la G	ly C	ys A		nr II 40	le Le	eu V	al Tl		er Le 45	eu A	la Me	

	Thr	A1a 50	Asp	Leu	Ile	Gly	Pro 55	His	Thr	Asn	Ser	Gly 60	Ala	Phe	Val	Tyr
5	Gly 65	Ser	Met	Ser	Phe	Leu 70	Asp	Lys	Val	Ala	Asn 75	Gly	Leu	Ala	Val	Met 80
10	Ala	Ile	Gln	Ser	Leu 85	His	Pro	Cys	Pro	Ser 90	Glu	Leu	Cys	Cys	Arg 95	Ala
- 0	Cys	Val	Ser	Phe 100	Tyr	His	Trp	Ala	Met 105	Val	Ala	Val	Thr	Gly 110	Gly	Val
15	Gly	Val	Ala 115	Ala	Ala	Leu	Cys	Leu 120	Суѕ	Ser	Leu	Leu	Leu 125	Trp	Pro	Thr
	Arg	Leu 130	Arg	Arg	Xaa											
20																
	(2)	INF	ORMA!	rion	FOR	SEQ	ID 1	VO : 2	230:							
25			(i) :	(A) L B) T D) T	ENGT: YPE : OPOL	H: 2 ami OGY:	8 am no a lin	ino cid ear	acid		: 230	O:			
30	Gly 1	Lys	Pro	Thr	Gly 5	Lys	Ser	Leu	Pro	Leu 10	Met	Trp	Met	Ile	Leu 15	Met
35	Gln	Pro	Ile	11e 20	Met	Ile	Ser	Met	Met 25	Ser	Asn	Gly				
	(2)	INFO	ORMAT	CION	FOR	SEQ	ID N	JO: 2	231:							
40			(i) :	()		ENGTI YPE :	H: 6 amin	1 am no a	ino a		S					
15			(xi)	SEQU	JENCE	E DES	SCRII	PTIO	N: SI	EQ II	ON C	231	l:			
	Met 1	Gln	Gly	Lys	Phe 5	Met	Lys	Val	Gln	Val 10	Tyr	Arg	Phe	Leu	Lys 15	Tyr
50	Leu	Leu	Met	Leu 20	Leu	Cys	Met	Phe	Val 25	Asn	Arg	Gly	Met	Ser 30	Lys	Asp
	Ser	Thr	Lys 35	Lys	Pro	Gly	Gln	Glu 40	Lys	Leu	Lys	Val	Ser 45	Leu	Gly	Ser
55	Ile	Leu 50	Asn	Met	Lys	Ser	Gln 55	Arg	Pro	Leu	Ser	Trp 60	Cys			
60	(2)	INFO	RMAT	NOI	FOR	SEQ	ID N	IO: 2	32:							

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```
(i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 29 amino acids
                    (B) TYPE: amino acid
5
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 232:
     Met Met Glu Arg Ser Met Met Ile Leu Leu Met Ala Ala Ser Met Thr
10
      Met Thr Ser Thr Gln Leu Trp Ser Phe Cys Cys Val His
                                       25
                   20
15
      (2) INFORMATION FOR SEQ ID NO: 233:
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 18 amino acids
                     (B) TYPE: amino acid
20
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 233:
      Met Trp Tyr Gln Leu Ala Lys Glu Glu Pro Gly Val Gly Ala Cys Ala
25
                                           10
      Leu Asp
30
      (2) INFORMATION FOR SEQ ID NO: 234:
              (i) SEQUENCE CHARACTERISTICS:
35
                     (A) LENGTH: 2 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 234:
40
       Leu Xaa
        1
       (2) INFORMATION FOR SEQ ID NO: 235:
45
              (i) SEQUENCE CHARACTERISTICS:
                      (A) LENGTH: 72 amino acids
                      (B) TYPE: amino acid
 50
                      (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 235:
       Met Leu Ile Cys Arg Leu Val Leu Leu Ala Asp Pro Gly Pro Val Asn
                                             10
 55
       Phe Met Val Arg Leu Phe Val Val Ile Val Met Phe Ala Trp Ser Ile
       Val Ala Ser Thr Ala Phe Leu Ala Asp Ser Gln Pro Pro Asn Arg Arg
 60
                                     40
```

	Ala	Leu 50	Ala	Val	Tyr	Pro	Val 55	Phe	Leu	Phe	Tyr	Phe 60	Val	Ile	Ser	Trp
5	Met 65	Ile	Leu	Thr	Phe	Thr 70	Pro	Gln								
10	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	236:							
15				(A) L B) T D) T	ENGT YPE : OPOL	H: 9 ami OGY:	6 am no a lin	ino cid ear	acid		: 23	6 :			
20	Met 1	Arg	Ser	Leu	Leu 5	Leu	Leu	Ser	Ala	Phe 10	Cys	Leu	Leu	Glu	Ala 15	Ala
	Leu	Ala	Ala	Glu 20	Val	Lys	Lys	Pro	Ala 25	Ala	Ala	Ala	Ala	Pro 30	Gly	Thr
25	Ala	Glu	Lys 35	Leu	Ser	Pro	Lys	Ala 40	Ala	Thr	Leu	Ala	Glu 45	Arg	Xaa	Pro
	Ala	Trp 50	Pro	Ser	Ala	Cys	Thr 55	Arg	Pro	Trp	Pro	Arg 60	Thr	Arg	Gln	Trp
30	Arg 65	Thr	Ser	Trp	Cys	His 70	Pro	Trp	Trp	Trp	Pro 75	Arg	Arg	Trp	Gly	Ser 80
35	Cys	Arg	Trp	Ala	Ala 85	Arg	Arg	Pro	Arg	Arg 90	Arg	Arg	Pro	Arg	Gln 95	Cys
40	(2)	INFO	ORMA:	rion	FOR	SEQ	ID 1	NO: 2	237 :							
1 5				(1	A) L 3) T O) T	ENGT YPE : OPOL	H: 1 ami: OGY:	43 ai no a line	mino cid ear	aci		: 237	7 :			
50	Met 1	Arg	Ser	Leu	Leu 5	Leu	Leu	Ser	Ala	Phe 10	Cys	Leu	Leu	Glu	Ala 15	Ala
	Leu	Ala	Ala	Glu 20	Val	Lys	Lys	Pro	Ala 25	Ala	Ala	Ala	Ala	Pro 30	Gly	Thr
55	Ala	Glu	Lys 35	Leu	Ser	Pro	Lys	Ala 40	Ala	Thr	Leu	Ala	Glu 45	Arg	Lys	Arg
60	Pro	Gly 50	Leu	Gln	Leu	Val	Pro 55	Gly	His	Gly	Gln	Gly 60	Pro	Gly	Ser	Gly

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	Glu His	s Pro	Gly	Val	Thr 70	Arg	Gly	Gly	Gly	Leu 75	Val	Ala	Gly	Ala	Arg 80
5	Val Ala	a Gly	Arg	Gln 85	Gly	Asp	His	Gly	Val 90	Ala	Gly	Gln	Gly	Ser 95	Ala
	Glu Ar	g Arg	Ala 100	Ala	Ala	Arg	Arg	Gly 105	Gly	Ala	Arg	Arg	Pro 110	Gly	Arg
10	Ala Al	a Ala 115	Leu	Thr	Gln	Gln	Leu 120	His	Gly	Ala	Gln	Arg 125	Asp	Leu	Glu
15	Ala Gl 13	_	Pro	Thr	Val	Arg 135	Thr	Gln	Leu	Ser	Glu 140	Leu	Arg	Xaa	
20	(2) IN		SEQU	ENCE (A) I (B) I (D) I	CHA ENGT YPE :	RACT H: 1 ami OGY:	ERIS .42 a .no a	TICS mind acid near	aci		22	0			
25	Met Ar 1				Leu				EEQ I Phe 10				Glu	Ala 15	Ala
30	Leu Al	la Ala	Glu 20		Lys	Lys	Pro	Ala 25		Ala	Ala	. Ala	Pro 30	Gly	Thr
	Ala G	lu Lys 35		ı Ser	Pro	Lys	Ala 40		Thr	Leu	Ala	Glu 45		Xaa	Arg
35	Pro G	ly Let 50	ı Glr	n Leu	ı Val	. Pro		/ His	s Gly	Gln	Gly 60		Gly	Ser	Gly
40	Glu H	is Pro	o Gly	y Val	Th:		g Gly	/ Gly	/ Gly	7 Leu 75		. Alā	a Gly	Ala	Arg 80
	Val A			85	5				90)				95	•
45	Glu A	rg Ar	g Ala 10		a Ala	a Arg	g Arg	g Gly 10!		/ Alá	a Arg	g Arq	110		/ Arg
	Ala A	la Al 11		u Thi	r Gli	n Gli	n Lei 120		a Gly	y Ala	a Gli	n Arg 12!) Lei	ı Glı
50	Ala G 1	ly G1 .30	n Pr	o Th	r Va	1 Arg		r Gl	n Lei	u Sei	r Gli 14		u Arg	ā	
55	(2) 1	NFORM													
60		(i)	SEÇ	(A)	LENC TYPE	STH: E: an	TERI 54 a nino 7: li	amino acio	aci i	.ds					

			(xi)	SEQ	UENC	E DE	SCRI	PTIO	N: S	EQ I	D NO	: 23	9 :			
5	Asp . 1	Pro	Glu	Ala	Ala 5	Asp	Ser	Gly	Glu	Pro 10	Gln	Asn	Lys	Arg	Thr 15	Pro
	Asp	Leu	Pro	Glu 20	Glu	Glu	Tyr	Val	Lys 25	Glu	Glu	Ile	Gln	Glu 30	Asn	Glu
10	Glu	Ala	Val 35	Lys	Lys	Met	Leu	Val 40	Glu	Ala	Thr	Arg	Glu 45	Phe	Glu	Glu
	Val	Val 50	Val	Asp	Glu	Ser										
15																
	(2)	INF		rion		_										
20				(A) L B) T D) T	ENGT YPE: OPOL	H: 6 ami OGY:	3 am no a lin	ino cid ear	acid		: 24	0 :			
25	Gln 1	Lys	Leu	Lys	Arg 5	Lys	Ala	Glu	Glu	Asp 10	Pro	Glu	Ala	Ala	Asp 15	Sei
30	Gly	Glu	Pro	Gln 20	Asn	Lys	Arg	Thr	Pro 25	Asp	Leu	Pro	Glu	Glu 30	Glu	Туг
	Val	Lys	Glu 35	Glu	Ile	Gln	Glu	Asn 40	Glu	Glu	Ala	Val	Lys 45	Lys	Met	Let
35	Val	Glu 50	Ala	Thr	Arg	Glu	Phe 55	Glu	Glu	Val	Val	Val 60	Asp	Glu	Ser	
40	(2)			(ENCE A) L B) T	CHAI ENGT YPE :	RACT H: 1	ERIS' 13 a no a	rics mino cid		ds					
45			(xi)	SEQI			OGY: SCRI			EQ I	D NO	: 24	1:			
	Lys 1	Ala	Met	Glu	Lys 5	Ser	Ser	Leu	Thr	Gln 10	His	Ser	Trp	Gln	Ser 15	Let
50	Lys	Asp	Arg	Tyr 20	Leu	Lys	His	Leu	Arg 25	Gly	Gln	Glu	His	Lys 30	Tyr	Leu
55	Leu	Gly	Asp 35	Ala	Pro	Val	Ser	Pro 40	Ser	Ser	Gln	Lys	Leu 45	Lys	Arg	Lys
	Ala	Glu 50	Glu	Asp	Pro	Glu	Ala 55	Ala	Asp	Ser	Gly	Glu 60	Pro	Gln	Asn	Lys
60	Arg 65	Thr	Pro	Asp	Leu	Pro 70	Glu	Glu	Glu	Tyr	Val 75	Lys	Glu	Glu	Ile	Glr 80

	Glu A	Asn (Glu	Glu	Ala 85	Val	Lys	Lys	Met :	Leu 90	Val	Glu	Ala	Thr .	Arg 95	Glu
5	Phe (Glu	Glu	Val 100	Val	Val	Asp	Glu	Ser 105	Pro	Pro	Asp	Phe	Glu 110	Ile	His
10	Ile															
10																
	(2)	INFC	RMA'	rion	FOR	SEQ	ID N	10: 2	:42:							
15			(i)	(A) L B) T	ENGT YPE :	H: 1 ami	ERIST 48 au no a lin	mino cid		ds					
20			(xi)	SEQ						EQ II	D NO	: 24	2 :			
20	Leu 1	Pro	Ser	Tyr	Asp 5	Glu	Ala	Glu	Arg	Thr 10	Lys	Ala	Glu	Ala	Thr 15	Ile
25	Pro	Leu	Val	Pro 20		Arg	Asp	Glu	Asp 25	Phe	Val	Gly	Arg	Asp 30	Asp	Phe
	Asp	Asp	Ala 35	Asp	Gln	Leu	Arg	Ile 40	Gly	Asn	Asp	Gly	Ile 45	Phe	Met	Leu
30	Thr	Phe 50		Met	Ala	Phe	Leu 55		Asn	Trp	Ile	Gly 60		Phe	Leu	Ser
25	Phe 65	Cys	Leu	ı Thr	Thr	Ser 70		Ala	Gly	Arg	Tyr 75		Ala	Ile	Ser	Gly 80
35	Phe	Gly	· Let	ı Ser	Leu 85		Lys	Trp	Ile	Leu 90		. Val	. Arg	Phe	Ser 95	Thr
40	Tyr	Ph∈	e Pro	o Gly 100		Phe	. Asr	Gly	Gln 105		Trp	Leu	Trp	Trp 110	Va]	l Phe
	Leu	Val	Le:	u Gly 5	/ Phe	e Leu	ı Lev	ı Ph∈ 120		Arg	σlΣ	Phe	2 Ile 125		тул	c Ala
45	Lys	Val		g Lys	s Met	. Pro	Gl:		Phe	e Sei	Ası	1 Lei 14(Arg	, Thi	r Arg
50	Val 145		ı Ph	e Ile	ę											
	(2)	IN	FORM	OITA	N FO	R SE(Q ID	NO:	243	:						
55			(i)	SEÇ	(A) (B)	LENG TYPE	TH: : an	24 a nino	mino acid	aci	.ds					
60			(x:	i) SE				: li RIPTI			ID N	10: 2	:43 :			

	Ala 1		Arg	Tyr	Gly 5		ıIl∈	e Ser	Gly	Ph∈		Leu	Ser	Leu	Il∈ 15	Lys
5	Ттр) Ile	e Leu	Ile 20		Arg	Phe	e Ser								
10	(2)	INF	ORMA	SEQU	ENCE	CHA	RACT	ERIS	TICS		ls					
15			(xi)	SEÇ			LOGY : SCRI			EQ I	:D N C): 24	4:			
	Met 1		His	Leu	Ser 5	Ala	Trp	Asn	Phe	Thr 10	Lys	Leu	Thr	Phe	Leu 15	Gln
20	Leu	Trp	Glu	Ile 20	Phe	Glu	Gly	Ser	Val 25	Glu	Asn	Cys	Gln	Thr 30	Leu	Thr
25			35	Lys	Leu	Gln	Ile	Lys 40	Tyr	Thr	Phe	Ser	Arg 45	Gly	Ser	Thr
	Pne	Tyr 50	He													
30	(2)	INF	ORMA'													
35				(A) L B) T D) T	ENGT YPE: OPOL	H: 2 ami OGY:	13 a no a lin	mino cid ear	aci		: 24	5:			
40	Phe 1	Ser	Ser	Asp	Phe 5	Arg	Thr	Ser	Pro	Trp	Glu	Ser	Arg	Arg	Val 15	Glu
	Ser	Lys	Ala	Thr 20	Ser	Ala	Arg	Cys	Gly 25	Leu	Trp	Gly	Ser	Gly 30	Pro	Arg
45	Arg	Arg	Pro 35	Ala	Ser	Gly	Met	Phe 40	Arg	Gly	Leu	Ser	Ser 45	Trp	Leu	Gly
50	Leu	Gln 50	Gln	Pro	Val	Ala	Gly 55	Gly	Gly	Gln	Pro	Asn 60	Gly	Asp	Ala	Pro
	Pro 65	Glu	Gln	Pro	Ser	Glu 70	Thr	Val	Ala	Glu	Ser 75	Ala	Glu	Glu	Glu	Leu 80
55			Ala		85					90					95	
	Asn	Tyr	Leu	Phe 100	Asn	Phe	Ala	Ser	Ala 105	Ala	Thr	Lys	Lys	Ile 110	Thr	Glu
60	Ser	Val	Ala	Glu	Thr	Ala	Gln	Thr	Ile	Lvs	Lvs	Ser	Val	Glu	Glu	Clu

			115					120					125			
5	Lys	Ile 130	Asp	Gly	Ile	Ile	Asp 135	Lys	Thr	Ile	Ile	Gly 140	Asp	Phe	Gln	Lys
5	Glu 145	Gln	Lys	Lys	Phe	Val 150	Glu	Glu	Gln	His	Thr 155	Lys	Lys	Ser	Glu	Ala 160
10	Ala	Val	Pro	Pro	Trp 165	Val	Asp	Thr	Asn	Asp 170	Glu	Glu	Thr	Ile	Gln 175	Gln
	Gln	Ile	Leu	Ala 180	Leu	Ser	Ala	Asp	Lys 185	Arg	Asn	Phe	Leu	Arg 190	Asp	Pro
15	Pro	Ala	Gly 195	Val	Gln	Phe	Asn	Phe 200	Asp	Phe	Asp	Gln	Met 205	Туr	Pro	Val
20	Ala	Leu 210	Val	Met	Leu											
	(2)	INF	ORMA	TION	FOR	SEQ	ID	NO:	246:							
25					(A) I (B) ' (D) '	LENG TYPE TOPO	rh: : am LOGY	49 ar ino a : lir	mino acid near	acio						
30) SEQ												
	Met		g Phe	e Ala	Lev S		l Pro	o Lys	: Leu	val 10		s Glu	ı Glu	ı Val	Phe 15	e Trp
35	Arg	g Ası	а Туз	r Phe		. Arg	g Vai	l Sei	Let 25		e Lys	s Glr	n Ser	Ala 30		n Leu
	Thi	r Ala	a Le		a Ala	a Gli	n Gl:	n Glr 40		a Ala	a Gly	y Ly:	s Gly 45		y Glı	u Glu
40	Glı	n														
45	(2) IN		OITA												
50				SEÇ i) SE	(A) (B) (D)	LENC TYPE TOPO	GTH: E: ar OLOG	76 a mino Y: li	mino acio inear	aci l		NO: 2	247:			
5 5	Se	er Th	ır Se	er Pr	:o G1	.y Vā 5	al Se	er Gl	u Ph		1 Se .0	er As	sp Al	a Ph		sp Ala .5
55	C7	/s As	sn Le		in Gl 20	ln Gl	lu As	sp Le		g Ly 25	rs Gl	lu Me	et Gl		n Le 30	eu Val
60	Le	eu As		ys Ly 35	/s G	ln G	lu G		nr Al 10	la Va	al L	eu Gi		lu As 15	sp Se	er Ala

```
Asp Trp Glu Lys Glu Leu Gln Glu Leu Gln Glu Tyr Glu Val Val
                               55
 5
      Thr Glu Ser Glu Lys Arg Asp Glu Asn Trp Asp Lys
                           70
10
      (2) INFORMATION FOR SEQ ID NO: 248:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 62 amino acids
                    (B) TYPE: amino acid
15
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 248:
      Ser Pro Trp Glu Ser Arg Arg Val Glu Ser Lys Ala Thr Ser Ala Arg
20
      Cys Gly Leu Trp Gly Ser Gly Pro Arg Arg Pro Ala Ser Gly Met
      Phe Arg Gly Leu Ser Ser Trp Leu Gly Leu Gln Gln Pro Val Ala Gly
25
                                  40
      Gly Gly Gln Pro Asn Gly Asp Ala Pro Pro Glu Gln Pro Ser
          50
30
      (2) INFORMATION FOR SEQ ID NO: 249:
             (i) SEQUENCE CHARACTERISTICS:
35
                    (A) LENGTH: 65 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 249:
40
      Pro Val Ala Gly Gly Gln Pro Asn Gly Asp Ala Pro Pro Glu Gln
                               10
      Pro Ser Glu Thr Val Ala Glu Ser Ala Glu Glu Glu Leu Gln Gln Ala
45
     Gly Asp Gln Glu Leu Leu His Gln Ala Lys Asp Phe Gly Asn Tyr Leu
                                  40
      Phe Asn Phe Ala Ser Ala Ala Thr Lys Lys Ile Thr Glu Ser Val Ala
50
                              55
     Glu
      65
55
      (2) INFORMATION FOR SEQ ID NO: 250:
             (i) SEQUENCE CHARACTERISTICS:
60
                    (A) LENGTH: 72 amino acids
```

```
(B) TYPE: amino acid
                   (D) TOPOLOGY: linear
            (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 250:
     Phe Gln Lys Glu Gln Lys Lys Phe Val Glu Glu Gln His Thr Lys Lys
 5
                                         10
       1
     Ser Glu Ala Ala Val Pro Pro Trp Val Asp Thr Asn Asp Glu Glu Thr
                                     25
10
     Ile Gln Gln Gln Ile Leu Ala Leu Ser Ala Asp Lys Arg Asn Phe Leu
                                 40
     Arg Asp Pro Pro Ala Gly Val Gln Phe Asn Phe Asp Phe Asp Gln Met
15
      Tyr Pro Val Ala Leu Val Met Leu
                          70
      65
20
      (2) INFORMATION FOR SEQ ID NO: 251:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 28 amino acids
25
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 251:
      Pro Phe Ile Cys Val Ala Arg Asn Pro Val Ser Arg Asn Phe Ser Ser
30
                              10
              5
      Pro Ile Leu Ala Arg Lys Leu Cys Glu Gly Ala Ala
                 20
35
       (2) INFORMATION FOR SEQ ID NO: 252:
             (i) SEQUENCE CHARACTERISTICS:
 40
                    (A) LENGTH: 33 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 252:
 45
       Lys Glu Asp Pro Ala Asn Thr Val Tyr Ser Thr Val Glu Ile Pro Lys
               5
                                          10
       Lys Met Glu Asn Pro His Ser Leu Leu Thr Met Pro Asp Thr Pro Arg
                                     25
 50
                   20
       Leu
 55
       (2) INFORMATION FOR SEQ ID NO: 253:
              (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 227 amino acids
 60
```

```
(B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 253:
      Ala Ser Ala Val Leu Leu Asp Leu Pro Asn Ser Gly Gly Glu Ala Gin
        1
      Ala Lys Lys Leu Gly Asn Asn Cys Val Phe Ala Pro Ala Asp Val Thr
                                      25
10
      Ser Glu Lys Asp Val Gln Thr Ala Leu Ala Leu Ala Lys Gly Lys Phe
                                  40
      Gly Arg Val Asp Val Ala Val Asn Cys Ala Gly Ile Ala Val Ala Ser
15
      Lys Thr Tyr Asn Leu Lys Lys Gly Gln Thr His Thr Leu Glu Asp Phe
20
     Gln Arg Val Leu Asp Val Asn Leu Met Gly Thr Phe Asn Val Ile Arg
                                          90
     Leu Val Ala Gly Glu Met Gly Gln Asn Glu Pro Asp Gln Gly Gly Gln
                                    105
25
      Arg Gly Val Ile Ile Asn Thr Ala Ser Val Ala Ala Phe Glu Gly Gln
     Val Gly Gln Ala Ala Tyr Ser Ala Ser Lys Gly Gly Ile Val Gly Met
30
                            135
     Thr Leu Pro Ile Ala Arg Asp Leu Ala Pro Ile Gly Ile Arg Val Met
                         150
                                            155
35
     Thr Ile Ala Pro Gly Leu Phe Gly Thr Pro Leu Leu Thr Ser Leu Pro
                                         170
     Glu Lys Val Cys Asn Phe Leu Ala Ser Gln Val Pro Phe Pro Ser Arg
                         185
40
     Leu Gly Asp Pro Ala Glu Tyr Ala His Leu Val Gln Ala Ile Ile Glu
     Asn Pro Phe Leu Asn Gly Glu Val Ile Arg Leu Asp Gly Ala Ile Arg
45
     Met Gln Pro
     225
50
     (2) INFORMATION FOR SEQ ID NO: 254:
             (i) SEQUENCE CHARACTERISTICS:
55
                    (A) LENGTH: 29 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 254:
60
     Ser Val Ala Ala Phe Glu Gly Gln Val Gly Gln Ala Ala Tyr Ser Ala
```

345

	1	5		10	15
5	Ser Lys	Gly Gly Ile Vai 20	l Gly Met Thr 25	Leu Pro Ile	Ala
	(2) INFO	RMATION FOR SE	Q ID NO: 255:		
10	·	(B) TYPE	TH: 61 amino : amino acid DLOGY: linear	acids	5:
15	Ala Arg	Arg Ser Gly Al	a Glu Leu Ala	Trp Asp Tyr	Leu Cys Arg Trp
20	Ala Gln	Lys His Lys As 20	n Trp Arg Phe 25	Gln Lys Thr	Arg Gln Thr Trp
	Leu Leu	Leu His Met Ty 35	r Asp Ser Asp 40	Lys Val Pro	Asp Glu His Phe 45
25	Ser Thr 50	Leu Leu Ala Ty	r Leu Glu Gly 55	Leu Gln Gly 60	Arg
30	(2) INFO	RMATION FOR SE	Q ID NO: 256:		
35		(B) TYPE	GTH: 22 amino E: amino acid DLOGY: linear	acids	ς.
33					Ser Gln Phe Tyr 15
40	Ile Asn	Lys Leu Cys Ph 20	ie		
45	(2) INFO	ORMATION FOR SE		S:	
50		(A) LEN (B) TYP	GTH: 22 amino E: amino acid OLOGY: linear	acids	57:
55	1	5 Asp Asn Ile G		Met Gln Asr 10	Ala Gln Leu Ser 15
		20			

BNSDOCID: <WO 9842738A1>

(2) INFORMATION FOR SEQ ID NO: 258:

```
(i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 25 amino acids
 5
                     (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 258:
      Lys Val Ser Tyr Leu Arg Pro Leu Asp Phe Glu Glu Ala Arg Glu Leu
10
                       5
      Phe Leu Leu Gly Gln His Tyr Val Phe
                   20
15
      (2) INFORMATION FOR SEQ ID NO: 259:
             (i) SEQUENCE CHARACTERISTICS:
20
                    (A) LENGTH: 25 amino acids
                     (E) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 259:
25
      Met Glu Arg Arg Cys Lys Met His Lys Arg Xaa Ile Ala Met Leu Glu
                                          10
      Pro Leu Thr Val Asp Leu Asn Pro Gln
                   20
30
      (2) INFORMATION FOR SEQ ID NO: 260:
35
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 23 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 260:
40
      Ser His Ile Val Lys Lys Ile Asn Asn Leu Asn Lys Ser Ala Leu Lys
      Tyr Tyr Gln Leu Phe Leu Asp
45
                   20
      (2) INFORMATION FOR SEQ ID NO: 261:
50
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 64 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
55
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 261:
      Phe Thr His Leu Ser Thr Cys Leu Leu Ser Leu Leu Leu Val Arg Met
                                           10
60
      Ser Gly Phe Leu Leu Leu Ala Arg Ala Ser Pro Ser Ile Cys Ala Leu
```

	20	25	30
	Asp Ser Ser Cys Phe Val	Gln Glu Tyr Cys Ser 40	Ser Tyr Ser Ser Ser 45
5	Cys Phe Leu His Gln His	Phe Pro Ser Leu Leu 55	Asp His Leu Cys Gln 60
10			
15	(B) TYPE	ARACTERISTICS: TH: 23 amino acids : amino acid	
20	, ,	LOGY: linear ESCRIPTION: SEQ ID NO	: 262:
	Phe Leu Leu Leu Ala Arg	g Ala Ser Pro Ser Ile 10	Cys Ala Leu Asp Ser 15
25	Ser Cys Phe Val Gln Glv 20	ı Tyr	
30	(2) INFORMATION FOR SEC		
35	(B) TYPE (D) TOPO	ARACTERISTICS: TH: 53 amino acids : amino acid DLOGY: linear ESCRIPTION: SEQ ID NO	v: 263:
40	Pro Asp Gly Arg Val Th 1 5 Phe Gly Met Ile Gly Le 20	10	15
45	Pro Gly Met Val His Le		
	Leu Asn Leu Asn Ser 50		
50			
55	(B) TYP		O: 264 :
60	Glu Asp Leu Leu Phe Ty	yr Leu Tyr Tyr Met Asr	n Gly Gly Asp Val Leu

	1				5					10					15	
5 .	Gln	Leu	Leu	Ala 20	Ala	Val	Glu	Leu	Phe 25	Asn	Arg	Asp	Trp	Arg 30	Tyr	His
	Lys	Glu	Glu 35	Arg	Val	Trp	Ile	Thr 40	Arg							
10	(2)	INF	ORMA'	rion	FOR	SEQ	ID I	NO:	265:							
15				(A) L B) T D) T	ENGT YPE: OPOL	H: 2 ami OGY:	4 am no a lin		acid		: 26	5:			
20	Val 1	His	Leu	Ala	Leu 5	Gly	Ser	Asp	Leu	Thr 10	Thr	Leu	Gly	Leu	Asn 15	Leu
	Asn	Ser	Pro	Glu 20	Asn	Leu	Tyr	Pro								
25																
	(2)	INF	ORMA	rion	FOR	SEQ	ID I	VO : 2	266:							
30			(i) (xi)	(A) L B) T D) T	ENGT: YPE: OPOL	H: 4 ami OGY:	l am no a lin	ear	acid		: 26	6 :			
35	His 1	Asn	Glu	Asp	Phe 5	Pro	Ala	Leu	Pro	Gly 10	Ser					
40	(2)	INFO	ORMA:	rion	FOR	SEQ	ID 1	VO: 2	267:							
			(i)	(.	A) L		H: 7	5 am	rics ino		s					
45			(xi)	(D) T	OPOL	OGY:	lin		EQ II	D NO	: 26	7:			
50	Gly 1	Arg	Ile	Ile	Asp 5	Thr	Ser	Leu	Thr	Arg 10	Asp	Pro	Leu	Val	Ile 15	Glu
	Leu	Gly	Gln	Lys 20	Gln	Val	Ile	P1	Gly 25	Leu	Glu	Gln	Ser	Leu 30	Leu	Asp
55	Met	Cys	Val 35	Gly	Glu	Lys	Arg	Arg 40	Ala	Ile	Ile	Pro	Ser 45	His	Leu	Ala
	Tyr	Gly 50	Lys ·	Arg	Gly	Phe	Pro 55	Pro	Ser	Val	Pro	Ala 60	Asp	Ala	Val	Val
60	Gln	Tyr	Asp	Val	Glu	Leu	Ile	Ala	Leu	Ile	Arg					

349

70 75 65 5 (2) INFORMATION FOR SEQ ID NO: 268: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 16 amino acids (B) TYPE: amino acid 10 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 268: Ile His Tyr Thr Gly Ser Leu Val Asp Gly Arg Ile Ile Asp Thr Ser 10 5 15 20 (2) INFORMATION FOR SEQ ID NO: 269: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 20 amino acids 25 (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 269: Cys Glu Ser Pro Glu Ser Pro Ala Gln Pro Ser Gly Ser Ser Leu Pro 30 Ala Trp Tyr His 35 (2) INFORMATION FOR SEQ ID NO: 270: (i) SEQUENCE CHARACTERISTICS: 40 (A) LENGTH: 95 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 270: 45 Glu Glu Ala Gly Ala Gly Arg Cys Ser His Gly Gly Ala Arg Pro Ala Gly Leu Gly Asn Glu Gly Leu Gly Leu Gly Gly Asp Pro Asp His 20 50 Thr Asp Thr Gly Ser Arg Ser Lys Gln Arg Ile Asn Asn Trp Lys Glu 40 Ser Lys His Lys Val Ile Met Ala Ser Ala Ser Ala Arg Gly Asn Gln 55 Asp Lys Asp Ala His Phe Pro Pro Pro Ser Lys Gln Ser Leu Leu Phe

Cys Pro Lys Ser Lys Leu His Ile His Arg Ala Glu Ile Ser Lys

350

95

90

85

(2) INFORMATION FOR SEQ ID NO: 271: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 23 amino acids (B) TYPE: amino acid 10 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 271: Ser Lys Gln Arg Ile Asn Asn Trp Lys Glu Ser Lys His Lys Val Ile 10 15 Met Ala Ser Ala Ser Ala Arg 20 20 (2) INFORMATION FOR SEQ ID NO: 272: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 32 amino acids 25 (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 272: Leu Phe His Trp Ala Cys Leu Asn Glu Arg Ala Ala Gln Leu Pro Arg 30 Asn Thr Ala Xaa Ala Gly Tyr Gln Cys Pro Ser Cys Asn Gly Pro Ser 25 35 40 (2) INFORMATION FOR SEQ ID NO: 273: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 185 amino acids (B) TYPE: amino acid 45 (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 273: Phe Tyr Ile Tyr Tyr Arg Pro Thr Asp Ser Asp Asn Asp Ser Asp Tyr 5 50 Lys Lys Asp Met Val Glu Gly Asp Lys Tyr Trp His Ser Ile Ser His Leu Gln Pro Glu Thr Ser Tyr Asp Ile Lys Met Gln Cys Phe Asn Glu 55 40 45 Gly Gly Glu Ser Glu Phe Ser Asn Val Met Ile Cys Glu Thr Lys Ala 60 Arg Lys Ser Ser Gly Gln Pro Gly Arg Leu Pro Pro Pro Thr Leu Ala

	65					70					75					80
-	Pro	Pro	Gln	Pro	Pro 85	Leu	Pro	Glu	Thr	Ile 90	Glu	Arg	Pro	Val	Gly 95	Thr
5	Gly	Ala	Met	Val 100	Ala	Arg	Ser	Ser	Asp 105	Leu	Pro	Tyr	Leu	Ile 110	Val	Gly
10	Val	Val	Leu 115	Gly	Ser	Ile	Val	Leu 120	Ile	Ile	Val	Thr	Phe 125	Ile	Pro	Phe
	Cys	Leu 130	Trp	Arg	Ala	Trp	Ser 135	Lys	Gln	Lys	His	Thr 140	Thr	Asp	Leu	Gly
15	Phe 145	Pro	Arg	Ser	Ala	Leu 150	Pro	Pro	Ser	Cys	Pro 155	Tyr	Thr	Met	Val	Pro 160
20	Leu	Gly	Gly	Leu	Pro 165	Gly	His	Gln	Ala	Val 170	Asp	Ser	Pro	Thr	Ser 175	Val
20	Ala	Ser	Val	Asp 180	Gly	Pro	Val	Leu	Met 185							
25	(2)	INF	ORMA'	rion	FOR	SEQ	ID	NO:	274:							
30				+	(A) I (B) 1 (D) 1	LENGT TYPE : TOPOI	H: 6 am: LOGY	56 an ino a : lir		acio): 27	4:			
35	Ту <i>г</i> 1	Ile	Tyr	Tyr	Arg		Thr	: Asp	Ser	Asp 10		Asp	Ser	. Asp	Туг 15	Lys
	Lys	Asp	Met	Val 20		Gly	Asp	Lys	Tyr 25		His	Ser	: Ile	Ser 30		: Leu
40	Gln	Pro	Glu 35		Ser	Tyr	Asp	11e		Met	Glr	ı Cys	Phe 45		ı Glu	l Gly
45	Gly	Glu 50		Glv	ı Phe	e Ser	Asr 55		. Met	: Ile	e Cys	Glu 60		Lys	s Ala	a Arg
43	Lys 65	Ser	c													
50	(2)	IN	FORM	OITA	v FOI	R SE(Q ID	NO:	275	:						
55				_	(A) (B) (D)	LENG TYPE TOPC	TH: : an LOGY	30 a nino : li	STIC mino acid near ON:	aci		o: 2	75 :			
		n Va	l Ar	g Al		u Le 5	u Hi	s Ar	g Me		o Gl O	u Pr	o Pr	o Ly	s Il 1	e Asn

	Thr	Ala	Lys	Phe 20	Asn	Asn	Asn	Lys	Arg 25	Lys	Asn	Leu	Ser	Leu 30		
5																
	(2)	INF	ORMA	rion	FOR	SEQ	ID i	NO: 2	276 :							
10			(i) (xi)	(A) L B) T D) T	CHA ENGT YPE: OPOL E DE	H: 1 ami OGY:	85 a no a lin	mino cid ear	aci		: 27	6 :			
15	Asn 1	Thr	Asn	Gln	Arg 5	Glu	Ala	Leu	Gln	Tyr 10	Ala	Lys	Asn	Phe	Gln 15	Pro
20	Phe	Ala	Leu	Asn 20	His	Gln	Lys	Asp	11e 25	Gln	Val	Leu	Met	Gly 30	Ser	Leu
	Val	Tyr	Leu 35	Arg	Gln	Gly	Ile	Glu 40	Asn	Ser	Pro	Tyr	Val 45	His	Leu	Leu
25	Asp	Ala 50	Asn	Gln	Trp	Ala	Asp 55	Ile	Cys	Asp	Ile	Phe 60	Thr	Arg	Asp	Ala
	Cys 65	Ala	Leu	Leu	Gly	Leu 70	Ser	Val	Glu	Ser	Pro 75	Leu	Ser	Val	Ser	Phe 80
30	Ser	Ala	Gly	Cys	Val 85	Ala	Leu	Pro	Ala	Leu 90	Ile	Asn	Ile	Lys	Ala 95	Val
35	Ile	Glu	Gln	Arg 100	Gln	Cys	Thr	Gly	Val 105	Trp	Asn	Gln	Lys	Asp 110	Glu	Leu
	Pro	Ile	Glu 115	Val	Asp	Leu	Gly	Lys 120	Lys	Cys	Trp	Tyr	His 125	Ser	Ile	Phe
40	Ala	Cys 130	Pro	Ile	Leu	Arg	Gln 135	Gln	Thr	Thr	Asp	Asn 140	Asn	Pro	Pro	Met
	Lys 145	Leu	Val	Cys	Gly	His 150	Ile	Ile	Ser	Arg	Asp 155	Ala	Leu	Asn	Lys	Met 160
45	Phe	Asn	Gly	Ser	Lys 165	Leu	Lys	Cys	Pro	Туг 170	Cys	Pro	Met	Glu	Gln 175	Ser
50	Pro	Gly	Asp	Ala 180	Lys	Gln	Ile	Phe	Phe 185							
	(2)	INF	ORMA!	rion	FOR	SEQ	ID I	NO: 2	277 :							
55			(i) :	(A) L B) T	CHATENGT YPE:	H: 6 ami	5 am no a	ino cid		s					
60			(xi)			OPOL E DE				EQ I	D NO	: 27	7 :			

	Ser 1	Tyr	Leu	Ser	Ala 5	Cys	Phe	Ala	Gly	Cys 10	Asn	Ser	Thr	Asn	Leu 15	Thr
5	Gly	Cys	Ala	Cys 20	Leu	Thr	Thr	Val	Pro 25	Ala	Glu	Asn	Ala	Thr 30	Val	Val
	Pro	Gly	Lys 35	Cys	Pro	Ser	Pro	Gly 40	Cys	Gln	Glu	Ala	Phe 45	Leu	Thr	Phe
10	Leu	Суs 50	Val	Met	Cys	Ile	Cys 55	Ser	Leu	Ile	Gly	Ala 60	Met	Ala	Arg	His
15	Pro 65															
20	(2)	INF	(i)	SEQU	ENCE (A) I (B) T	SEQ CHA LENGT TYPE: TOPOL CE DE	RACT H: 8 ami	ERIS 34 am ino a lir	TICS uino cid ear	acid		: 27	8:			
25	Pro 1					Leu					Ser			Leu	Lys 15	Ser
30	Туг	· Ala	. Lev	Gly 20		. Leu	Phe	e Leu	Leu 25		ı Arg	Leu	. Leu	Gly 30		Ile
	Pro	Pro	Pro 35		ı Ile	e Phe	: Gly	Ala 40		′ Il∈	e Asp	Ser	Thr 45		Leu	Phe
35	Trp	Ser 50		r Ph∈	e Cys	s Gly	Glu 55		n Gly	Alā	a Cys	Val		Tyr	Asp	Asn
40	65	5		r Arg		Leu 70		r Val	l Ser	: Ile	e Ala 75		e Ala	ı Lev	ı Lys	80 80
45	(2) IN	FORM	ATIO	N FO	R SE(Q ID	NO:	279	:						
50					(A) (B) (D)	E CH LENG TYPE TOPC	TH: : am LOGY	182 nino 7: li	amin acid near	o ac		0: 2	79:			
55	Gl	n Se 1	r Le	eu Ph	e Th	r Ar	g Ph	e Va	l Ar		1 G1 0	y Va	l Pr	o Th	r Vai	l Asp 5
	Le	eu As	p Al		n Gl 0	y Ar	g Al	a Ar		a Se 5	r Le	u Cy	s Xa	a Xa 3		r Asn
60	Tr	p Ar	g Ty	r Ly	rs As	n Le	u Gl	y As	n Le	u Pr	o Hi	s Va	1 G1	n Le	u Le	u Pro

			35					40					45			
5	Glu	Phe 50	Ser	Thr	Ala	Asn	Ala 55	Gly	Leu	Leu	Tyr	Asp 60	Phe	Gln	Leu	Ile
J	Asn 65	Val	Glu	Asp	Phe	Gln 70	Gly	Val	Gly	Glu	Ser 75	Glu	Pro	Asn	Pro	Туг 80
10	Phe	Tyr	Gln	Asn	Leu 85	Gly	Glu	Ala	Glu	Tyr 90	Val	Val	Ala	Leu	Phe 95	Met
	Tyr	Met	Cys	Leu 100	Leu	Gly	Tyr	Pro	Ala 105	Asp	Lys	Ile	Ser	Ile 110	Leu	Thr
15	Thr	Tyr	Asn 115	Gly	Gln	Lys	His	Leu 120	Ile	Arg	Asp	Ile	Ile 125	Asn	Arg	Arg
20	Cys	Gly 130	Asn	Asn	Pro	Leu	Ile 135	Gly	Arg	Pro	Asn	Lys 140	Val	Thr	Thr	Val
20	Asp 145	Arg	Phe	Gln	Gly	Gln 150	Gln	Asn	Asp	Tyr	Ile 155	Leu	Leu	Ser	Leu	Val 160
25	Arg	Thr	Arg	Ala	Val 165	Gly	His	Leu	Arg	Asp 170	Val	Arg	Arg	Leu	Val 175	Val
	Ala	Met	Ser	Arg 180	Ala	Arg										
30																
	(2)	INF	ORMA	NOI	FOR	SEQ	ID 1	10: 2	280:							
35				(A) L B) T D) T	ENGT: YPE: OPOL	H: 7 ami OGY:	ERIST 7 am no a lin PTIO	ino . cid ear	acid		: 28	O:			
40	Leu 1	Val	Lys	Glu	Ala 5	Lys	Ile	Ile	Ala	Met 10	Thr	Cys	Thr	His	Ala 15	Ala
45	Leu	Lys	Arg	His 20	Asp	Leu	Val	Lys	Leu 25	Gly	Phe	Lys	Tyr	Asp 30	Asn	Ile
43	Leu	Met	Glu 35	Glu	Ala	Ala	Gln	Ile 40	Leu	Glu	Ile	Glu	Thr 45	Phe	Ile	Pro
50	Leu	Leu 50	Leu	Gln	Asn	Pro	Gln 55	Asp	Gly	Phe	Ser	Arg 60	Leu	Lys	Arg	Trp
	Ile 65	Met	Ile	Gly	Asp	His 70	His	Gln	Leu	Pro	Pro 75	Val	Ile			
55																
	(2)	INF	ORMA!	NOI	FOR	SEQ	ID I	NO: 2	281:							
60			(i)					ERIS 25 a			ds					

	(B) TYPE: amino acid (D) TOPOLOGY: linear															
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 281:															
5	Asp 1	Thr	Tyr	Pro	Asn 5	Glu	Glu	Lys	Gln	Gln 10	Glu	Arg	Val	Phe	Pro 15	Xaa
10	Xaa	Ser	Ala	Met 20	Val	Asn	Asn	Gly	Ser 25	Leu	Ser	Tyr	Asp	His 30	Glu	Arg
10	Asp	Gly	Arg 35	Pro	Thr	Glu	Leu	Gly 40	Gly	Cys	Xaa	Ala	Ile 45	Val	Arg	Asn
15	Leu	His 50	Tyr	Asp	Thr	Phe	Leu 55	Val	Ile	Arg	Tyr	Val 60	Lys	Arg	His	Leu
	Thr 65	Ile	Met	Met	Asp	Ile 70	Asp	Gly	Lys	His	Glu 75	Trp	Arg	Asp	Cys	Ile 80
20	Glu	Val	Pro	Gly	Val 85	Arg	Leu	Pro	Arg	Gly 90	Tyr	Tyr	Phe	Gly	Thr 95	Ser
25	Ser	Ile	Thr	Gly 100	Asp	Leu	Ser	Asp	Asn 105	His	Asp	Val	Ile	Ser 110	Leu	Lys
23	Leu	Phe	Glu 115	Leu	Thr	Val	Glu	Arg 120	Thr	Pro	Glu	Glu	Glu 125			
30	(2)	INF	'ORMA'													
35					(A) I (B) T (D) T	ENGI TYPE : TOPOI	TH: 8 am: LOGY	35 ar ino a : lir	mino acid near	acio	ds ID NO): 28	32:			
40	Leu 1		Arg	Glu	His		Leu	Ser	Lys	Pro		Glr	Gly	Val	Gly 15	Thr
	Gly	Sei	s Ser	Ser 20		Trp) Asr	Leu	Met 25		y Asr	ı Alā	Met	Val		Thr
45	Gln	ту	r Ile 35		, Leu	Thr	Pro	Asp 40		: Glr	n Ser	Lys	Glr 45		/ Ala	a Leu
50	Trp	Ası 5		g Val	L Pro	Cys	s Phe 5		ı Ar	g Ası	p Trp	Gl: 60		ı Glr	ı Val	l His
50	Phe 65		s Ile	e His	s Gly	Glr 70		/ Ly:	s Ly:	s As:	n Lei 7!		s Gly	y Ası	Gly	y Leu 80
55	Ala	a Il	e Trī	о Ту	r Thi											
60	(2)) IN	FORM	ATIO	N FO	R SE	Q ID	NO:	283	:						

```
(i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 32 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
 5
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 283:
      Pro Gly Thr Leu Gln Cys Ser Ala Leu His His Asp Pro Gly Cys Ala
                        5
10
      Asn Cys Ser Arg Phe Cys Arg Asp Cys Ser Pro Pro Ala Cys Gln Cys
                                       25
15
      (2) INFORMATION FOR SEQ ID NO: 284:
20
              (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 27 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 284:
25
      Phe Leu Tyr Asp Val Leu Met Xaa His Glu Ala Val Met Arg Thr His
                                           10
      Gln Ile Gln Leu Pro Asp Pro Glu Phe Pro Ser
30
                   20
      (2) INFORMATION FOR SEQ ID NO: 285:
35
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 6 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
40
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 285:
      Gly Trp Tyr Trp Cys Gly
45
      (2) INFORMATION FOR SEQ ID NO: 286:
             (i) SEQUENCE CHARACTERISTICS:
50
                     (A) LENGTH: 129 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 286:
55
      Met Lys Val Gly Ala Arg Ile Arg Val Lys Met Ser Val Asn Lys Ala
      His Pro Val Val Ser Thr His Trp Arg Trp Pro Ala Glu Trp Pro Gln
                                       25
                                                           30
60
```

	Met	Phe	Leu 35	His	Leu	Ala	Gln	Glu 40	Pro	Arg	Thr	Glu	Val 45	Lys	Ser	Arg
5	Pro	Leu 50	Gly	Leu	Ala	Gly	Phe 55	Ile	Arg	Gln	Asp	Ser 60	Lys	Thr	Arg	Lys
	Pro 65	Leu	Glu	Gln	Glu	Thr 70	Ile	Met	Ser	Ala	Ala 75	Asp	Thr	Ala	Leu	Trp 80
10	Pro	Tyr	Gly	His	Gly 85	Asn	Arg	Glu	His	Gln 90	Glu	Asn	Glu	Leu	Gln 95	Lys
15	Tyr	Leu	Gln	Туr 100	Lys	Asp	Met	His	Leu 105	Leu	Asp	Ser	Gly	Gln 110	Ser	Leu
13	Gly	His	Thr 115		Thr	Leu	Gln	Gly 120		His	Asn	Leu	Thr 125	Ala	Leu	Asn
20	Ile															
25	(2)	INF	ORMA	AOIT!	I FOF	R SEÇ	Į ID	NO:	287:							
23			(i)		(A)	LENG'	TH:	reris 49 au ino	mino		ds					
30			(xi		QUEN	CE DI	ESCR	: li:	: NC							
		c Lei l	ı His	s Ly:		n Sei	r Vai	l Sei	c Glr	n Ile 10		r Val	l Lev	ı Sei	6 Gly 19	y Gly 5
35				2	0				2!	5				3(0	y Met
40	Se	r Il	e Tr		p Va	l Ly	s Se	r Le 4		u Se:	r Al	a Le	u Ly: 4	s As _l 5	o Le	u Lys
	Il	е														
45	(2) IN	IFORM	L ATIC	N FC	R SE	Q II	NO:	288	:						
50			(i)	SEÇ	(A)	LEN	GTH:	CTERI 21 a mino	amino	ac:	ids					
			(x:	i) SI	(D)	TOP	OLOG	Y: 1	inea	r	ID 1	NO: 2	288:			
55	G]	lu Al 1	la Se	er Ly	/s Se	er Se 5	er H	is Al	la G]		eu As LO	sp Le	eu Ph	ne Se		al Ala 15
	A	la C <u>r</u>	ys H:	is Aı 2	rg Pl 20	ne										
60																

	(2) INFORMATION FOR SEQ ID NO: 289:
5	(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 21 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 289:
10	Tyr Met Gly Lys Gly Ser Met Thr Gly Leu Ala Leu Lys His Met Phe 1 5 10 15
15	Glu Arg Ser Phe Thr 20
20	(2) INFORMATION FOR SEQ ID NO: 290: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 27 amino acids (B) TYPE: amino acid
25	(D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 290: Val Thr Gly Ile Ile Asp Ser Leu Thr Ile Ser Pro Lys Ala Ala Arg 1 5 10 15
30	Val Gly Leu Gln Tyr Ser Thr Gln Val His 20 25
35	(2) INFORMATION FOR SEQ ID NO: 291: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 24 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear
40	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 291: Thr Glu Phe Thr Leu Arg Asn Phe Asn Ser Ala Lys Asp Met Lys Lys 1 10 15
45	1 5 10 15 Ala Val Ala His Met Lys Tyr Met 20
50	(2) INFORMATION FOR SEQ ID NO: 292: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 27 amino acids
55	(B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 292:
60	Gly Lys Gly Ser Met Thr Gly Leu Ala Leu Lys His Met Phe Glu Arg 1 5 10 15

Ser Phe Thr Gln Gly Glu Gly Ala Arg Pro Phe 5 (2) INFORMATION FOR SEQ ID NO: 293: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 44 amino acids 10 (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 293: Ser Thr Arg Val Pro Arg Ala Ala Ile Val Phe Thr Asp Gly Arg Ala 15 Gln Asp Asp Val Ser Glu Trp Ala Ser Lys Ala Lys Ala Asn Gly Ile 25 Thr Met Tyr Ala Val Gly Val Gly Lys Ala Ile Glu 20 40 35 25 (2) INFORMATION FOR SEQ ID NO: 294: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 42 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear 30 (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 294: Glu Glu Leu Gln Glu Ile Ala Ser Glu Pro Thr Asn Lys His Leu Phe 35 Tyr Ala Glu Asp Phe Ser Thr Met Asp Glu Ile Ser Glu Lys Leu Lys 25 Lys Gly Ile Cys Glu Ala Leu Glu Asp Ser 40 35 (2) INFORMATION FOR SEQ ID NO: 295: 45 (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 11 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 295: 50 Thr Gln Arg Leu Glu Glu Met Thr Gln Arg Met 1 5 55 (2) INFORMATION FOR SEQ ID NO: 296:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 10 amino acids

```
(B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 296:
      Pro Gln Gly Cys Pro Glu Gln Pro Leu His
10
      (2) INFORMATION FOR SEQ ID NO: 297:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 33 amino acids
                     (B) TYPE: amino acid
15
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 297:
      Arg Cys Lys Lys Cys Thr Glu Gly Pro Ile Asp Leu Val Phe Val Ile
                                           10
20
      Asp Gly Ser Lys Ser Leu Gly Glu Glu Asn Phe Glu Val Val Lys Gln
                  20
                                       25
      Phe
25
      (2) INFORMATION FOR SEQ ID NO: 298:
30
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 60 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
35
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 298:
      Met Ala Ala Leu Leu Arg His Val Gly Arg His Cys Leu Arg Ala
40
      His Phe Ser Pro Gln Leu Cys Ile Arg Asn Ala Val Pro Leu Gly Thr
      Thr Ala Lys Glu Glu Met Glu Arg Phe Trp Asn Lys Asn Ile Gly Ser
45
      Asn Arg Pro Leu Ser Pro His Ile Thr Ile Tyr Ser
           50
50
      (2) INFORMATION FOR SEQ ID NO: 299:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 32 amino acids
55
                     (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 299:
      Val Phe Pro Leu Met Tyr His Thr Trp Asn Gly Ile Arg His Leu Met
60
                                           10
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```
Trp Asp Leu Gly Lys Gly Leu Lys Ile Pro Gln Leu Tyr Gln Ser Gly
                                     25
5
10
     (2) INFORMATION FOR SEQ ID NO: 300:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 17 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
15
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 300:
     Met Ala Ala Leu Leu Leu Arg His Val Gly Arg His Cys Leu Arg Ala
                                      10
                      5
20
      His
25
      (2) INFORMATION FOR SEQ ID NO: 301:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 18 amino acids
30
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 301:
      Val Lys Ser Leu Cys Leu Gly Pro Ala Leu Ile His Thr Ala Lys Phe
35
                                   10
      Ala Leu
40
      (2) INFORMATION FOR SEQ ID NO: 302:
             (i) SEQUENCE CHARACTERISTICS:
45
                    (A) LENGTH: 23 amino acids
                     (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 302:
      Val Phe Pro Leu Met Tyr His Thr Trp Asn Gly Ile Arg His Leu Met
 50
                                          10
              5
        1
      Trp Asp Leu Gly Lys Gly Leu
                   20
 55
       (2) INFORMATION FOR SEQ ID NO: 303:
 60
             (i) SEQUENCE CHARACTERISTICS:
```

```
(A) LENGTH: 22 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 303:
 5
      Arg Val Trp Asp Val Arg Pro Phe Ala Pro Lys Glu Arg Cys Val Lys
                                          10
      Ile Phe Gln Gly Asn Val
10
      (2) INFORMATION FOR SEQ ID NO: 304:
15
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 30 amino acids
                     (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
20
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 304:
      His Asn Phe Glu Lys Asn Leu Leu Arg Cys Ser Trp Ser Pro Asp Gly
25
      Ser Lys Ile Ala Ala Gly Ser Ala Asp Arg Phe Val Tyr Val
                   20
                                      25
30
      (2) INFORMATION FOR SEQ ID NO: 305:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 30 amino acids
                     (B) TYPE: amino acid
35
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 305:
      Trp Asp Thr Thr Ser Arg Arg Ile Leu Tyr Lys Leu Pro Gly His Ala
                                  10
40
      Gly Ser Ile Asn Glu Val Ala Phe His Pro Asp Glu Pro Ile
45
      (2) INFORMATION FOR SEQ ID NO: 306:
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 20 amino acids
50
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 306:
      Val Arg Gly Arg Thr Val Leu Arg Pro Gly Leu Asp Ala Glu Pro Glu
55
       1
                       5
                                           10
      Leu Ser Pro Glu
                   20
60
```

```
(2) INFORMATION FOR SEQ ID NO: 307:
             (i) SEQUENCE CHARACTERISTICS:
5
                    (A) LENGTH: 19 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 307:
      Glu Gln Arg Val Leu Glu Arg Lys Leu Lys Lys Glu Arg Lys Lys Glu
10
                                           10
      Glu Arg Gln
15
      (2) INFORMATION FOR SEQ ID NO: 308:
20
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 13 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 308:
25
      Arg Leu Arg Glu Ala Gly Leu Val Ala Gln His Pro Pro
30
       (2) INFORMATION FOR SEQ ID NO: 309:
              (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 17 amino acids
35
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 309:
      Gly Arg Ile Pro Ala Pro Ala Pro Ser Val Pro Ala Gly Pro Asp Ser
40
                        5
       Arg
45
       (2) INFORMATION FOR SEQ ID NO: 310:
              (i) SEQUENCE CHARACTERISTICS:
 50
                     (A) LENGTH: 42 amino acids
                      (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 310:
       Thr Gly Cys Val Leu Val Leu Ser Arg Asn Phe Val Gln Tyr Ala Cys
 55
         1
       Phe Gly Leu Phe Gly Ile Ile Ala Leu Gln Thr Ile Ala Tyr Ser Ile
                                        25
 60
```

Leu Trp Asp Leu Lys Phe Leu Met Arg Asn

```
35
 5
      (2) INFORMATION FOR SEQ ID NO: 311:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 55 amino acids
10
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 311:
      Ser Arg Ser Glu Gly Lys Ser Met Phe Ala Gly Val Pro Thr Met Arg
15
      Glu Ser Ser Pro Lys Gln Tyr Met Gln Leu Gly Gly Arg Val Leu Leu
                                       25
20
      Val Leu Met Phe Met Thr Leu Leu His Phe Asp Ala Ser Phe Phe Ser
      Ile Val Gln Asn Ile Val Gly
           50
25
      (2) INFORMATION FOR SEQ ID NO: 312:
30
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 60 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 312:
35
      Gly Thr Ala Glu Asp Phe Ala Asp Gln Phe Leu Arg Val Thr Lys Gln
                                          10
      Tyr Leu Pro His Val Ala Arg Leu Cys Leu Ile Ser Thr Phe Leu Glu
40
      Asp Gly Ile Arg Met Trp Phe Gln Trp Ser Glu Gln Arg Asp Tyr Ile
45
      Asp Thr Trp Asn Cys Gly Tyr Leu Leu Ala Ser
50
      (2) INFORMATION FOR SEQ ID NO: 313:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 17 amino acids
                    (B) TYPE: amino acid
55
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 313:
      Ala Ser Phe Leu Leu Ser Arg Thr Ser Trp Gly Thr Ala Leu Met Ile
                       5
                                           10
60
```

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365

Leu

```
5
     (2) INFORMATION FOR SEQ ID NO: 314:
            (i) SEQUENCE CHARACTERISTICS:
                   (A) LENGTH: 8 amino acids
10
                    (B) TYPE: amino acid
                   (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 314:
     Leu Met Arg Asn Glu Ser Arg Ser
15
                      5
      1
      (2) INFORMATION FOR SEQ ID NO: 315:
20
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 13 amino acids
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 315:
25
      Ala Ser Phe Leu Leu Ser Arg Thr Ser Trp Gly Thr Ala
       1 5
                                         10
30
      (2) INFORMATION FOR SEQ ID NO: 316:
             (i) SEQUENCE CHARACTERISTICS:
                   (A) LENGTH: 20 amino acids
35
                    (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 316:
      Phe Ile Ser Phe Ala Asn Ser Arg Ser Ser Glu Asp Thr Lys Gln Met
 40
                             10
      Met Ser Ser Phe
 45
       (2) INFORMATION FOR SEQ ID NO: 317:
 50
              (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 27 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
              (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 317:
 55
       Asp Pro Arg Arg Pro Asn Lys Val Leu Arg Tyr Lys Pro Pro Pro Ser
              5
        1
       Glu Cys Asn Pro Ala Leu Asp Asp Pro Thr Pro
 60
```

```
(2) INFORMATION FOR SEQ ID NO: 318:
 5
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 30 amino acids
                     (B) TYPE: amino acid
                     (D) TOPOLOGY: linear
10
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 318:
      Asp Tyr Met Asn Leu Leu Gly Met Ile Phe Ser Met Cys Gly Leu Met
                                           10
15
      Leu Lys Leu Lys Trp Cys Ala Trp Val Ala Val Tyr Cys Ser
                   20
                                       25
20
      (2) INFORMATION FOR SEQ ID NO: 319:
             (i) SEQUENCE CHARACTERISTICS:
                     (A) LENGTH: 22 amino acids
                     (B) TYPE: amino acid
25
                     (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 319:
      Met Leu Ser Ile Ser Ala Val Val Met Ser Tyr Leu Gln Asn Pro Gln
30
      Pro Met Thr Pro Pro Trp
                   20
35
      (2) INFORMATION FOR SEQ ID NO: 320:
             (i) SEQUENCE CHARACTERISTICS:
                    (A) LENGTH: 52 amino acids
40
                     (B) TYPE: amino acid
                    (D) TOPOLOGY: linear
             (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 320:
      Ala Ala Gly Asp Gly Asp Val Lys Leu Gly Thr Leu Gly Ser Gly Ser
45
                                      10
      Glu Ser Ser Asn Asp Gly Gly Ser Glu Ser Pro Gly Asp Ala Gly Ala
50
      Ala Ala Xaa Gly Gly Gly Trp Ala Ala Ala Ala Leu Ala Leu Leu Thr
                                   40
      Gly Gly Glu
           50
55
      (2) INFORMATION FOR SEQ ID NO: 321:
60
             (i) SEQUENCE CHARACTERISTICS:
```

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	(A) LENGTH: 177 amino acids(B) TYPE: amino acid(D) TOPOLOGY: linear															
5			(xi)		-					EQ II	ON C	: 321	.:			
J	Ala 1	Ala	Asp	Asn	Tyr 5	Gly	Ile	Pro	Arg	Ala 10	Cys	Arg	Asn	Ser	Ala 15	Arg
10	Ser	Tyr	Gly	Ala 20	Ala	Trp	Leu	Leu	Leu 25	Xaa	Pro	Ala	Gly	Ser 30	Ser	Arg
	Val	Glu	Pro 35	Thr	Gln	Asp	Ile	Ser 40	Ile	Ser	qzA	Gln	Leu 45	Gly	Gly	Gln
15	Asp	Va1 50	Pro	Val	Phe	Arg	Asn 55	Leu	Ser	Leu	Leu	Val 60	Val	Gly	Val	Gly
20	Ala 65	Val	Phe	Ser	Leu	Leu 70	Phe	His	Leu	Gly	Thr 75	Arg	Glu	Arg	Arg	Arg 80
_~	Pro	His	Ala	Xaa	Glu 85	Pro	Gly	Glu	His	Thr 90	Pro	Leu	Leu	Ala	Pro 95	Ala
25	Thr	Ala	Gln	Pro 100	Leu	Leu	Leu	Trp	Lys 105	His	Trp	Leu	Arg	Glu 110	Xaa	Ala
	Phe	Tyr	Gln 115	Val	Gly	Ile	Leu	Tyr 120	Met	Thr	Thr	Arg	Leu 125	Ile	Val	Asn
30	Leu	Ser 130	Gln	Thr	Tyr	Met	Ala 135		Tyr	Leu	Thr	Tyr 140	Ser	Leu	His	Leu
35	Pro 145	_	Lys	Phe	Ile	Ala 150	Thr	Ile	Pro	Leu	Val 155	Met	Tyr	Leu	Ser	Gly 160
	Phe	Leu	Ser	Ser	Phe 165	Leu	Met	Lys	Pro	Ile 170		Lys	Cys	Ile	Gly 175	Arg
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	Ser	Le	1 Lys 35		Lys	s Lys	s Asp	Ser 40		r Gly	/ Alā	Pro	Ser 45		Pro	Ile
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10	Phe	Pro	Ala	Pro 100	Pro	Lys	Gln	Leu	Asp 105	Met	Gly	Asp	Glu	Val 110	Tyr	Asp
	Asp	Val	Asp 115	Thr	Ser	Asp	Phe	Pro 120	Val	Ser	Ser	Ala	Glu 125	Met	Ser	Gln
15	Gly	Thr 130	Asn	Val	Gly	Lys	Ala 135	Lys	Thr	Glu	Glu	Lys 140	Asp	Leu	Lys	Lys
20	Leu 145	Lys	Lys	Gln	Xaa	Lys 150	Glu	Xaa	Lys	Asp	Phe 155	Arg	Lys	Lys	Phe	Lys 160
20	Тут	Asp	Gly	Glu	Ile 165	Arg	Val	Leu	Tyr	Ser 170	Thr	Lys	Val	Thr	Thr 175	Ser
25	Ile	Thr	Ser	Lys 180	Lys	Trp	Gly	Thr	Arg 185	Asp	Leu	Gln	Val	Lys 190	Pro	Gly
	Glu	Ser	Leu 195	Glu	Val	Ile	Gln	Thr 200	Thr	Asp	Asp	Thr	Lys 205	Val	Leu	Cys
30	Arg	Asn 210	Glu	Glu	Gly	Lys	Tyr 215	Gly	Tyr	Val	Leu	Arg 220	Ser	Tyr	Leu	Ala
35	Asp 225		Asp	Gly	Glu	Ile 230	Tyr	Asp	Asp	Ile	Ala 235	Asp	Gly	Cys	Ile	Туг 240
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55	Arg	His	Ala 35		Gly	Gly	Val	His 40	Ile	Glu	Pro	Arg	Туr 45	Arg	Gln	Phe
	Pro	Gln 50		Thr	Arg	Ser	Gln 55	Val	Phe	Gln	Ser	Glu 60	Phe	Phe	Ser	Gly
60	Leu	Met	Tro	Phe	Tro	Ile	Leu	Trp	Ara	Phe	Tro	His	Asp	Ser	Glu	Glı

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Val Leu Gly His Phe Pro Tyr Pro Asp Pro Ser Gln Trp Thr Asp Glu 85 90 95

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INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

(PCT Rule 13bis)

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3. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet
Name of depositary institution American T	Type Culture Collection
American	spe Cunute Conection
Address of depositary institution (including posi-	tal code and country)
10801 University Boulevard Manassas, Virginia 20110-2209 United States of America	
Date of deposit March 7, 1997	Accession Number 97923
C. ADDITIONAL INDICATIONS (leave t	blank if not applicable) This information is continued on an additional sheet
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INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

(PCT Rule 13bis)

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3. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet
Name of depositary institution American Type Cultu	ire Collection
Address of depositary institution (including postal code and 10801 University Boulevard Manassas, Virginia 20110-2209 United States of America	l country)
Date of deposit May 22, 1997	Accession Number 209071
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INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

(PCT Rule 13bis)

Address of depositary institution (including postal code and country) 10801 University Boulevard Manassas. Virginia 20110-2209 United States of America Date of deposit February 25, 1998 Accession Number 209641 C. ADDITIONAL INDICATIONS (leave blank if not applicable) This information is continued on an additional sheet D. DESIGNATED STATES FOR WHICH INDICATIONS ARE MADE (if the indications are not for all designated State) E. SEPARATE FURNISHING OF INDICATIONS (leave blank if not applicable) The indications listed below will be submitted to the International Bureau later (species the separal square at the indications as a location of the indications is continued on the indication of the indicatio	A. The indications made below relate to the mon page 73	line N/A
American Type Culture Collection Address of depositary institution (including postal code and country) 10801 University Boulevard Manassas. Virginia 20110-2209 United States of America Date of deposit February 25, 1998 Accession Number 209641 C. ADDITIONAL INDICATIONS (leave blank if not applicable) This information is continued on an additional sheet D. DESIGNATED STATES FOR WHICH INDICATIONS ARE MADE (if the indications are not for all designated State) E. SEPARATE FURNISHING OF INDICATIONS (leave blank if not applicable) The indications listed below will be submitted to the International Bureau later (specify the general nature of the indications, e.g., "(cees Number of Deposit) For receiving Office use only For International Bureau use only This sheet was received with the international application Authorized officer	B. IDENTIFICATION OF DEPOSIT	Further descriptions identified an actual invest shows
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Applicant's or agent's file Z004PCT reference number	International application Unassigned

INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

(PCT Rule 13bis)

A. The indications ma	ade below relate to the microorgan	nism referred to in the description ine N/A
B. IDENTIFICATI	ON OF DEPOSIT	Further deposits are identified on an additional sheet
Name of depositary ins	American Type Cu	ulture Collection
Address of depositary 10801 University B Manassas, Virginia United States of Am	20110-2209	e and country)
Date of deposit July	y 24, 1997	Accession Number 209179
C. ADDITIONAL	INDICATIONS (leave blank if n	Inot applicable) This information is continued on an additional sheet
D. DESIGNATED	STATES FOR WRICH IND	DICATIONS ARE MADE (if the indications are not for all designated States)
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(PCT Rule 13bis)

on page 77	ade below relate to the microorg	anism referred to	o in the description
3. IDENTIFICAT	ION OF DEPOSIT		Further deposits are identified on an additional sheet
Name of depositary in	American Type (Culture Collec	tion
Address of depositary	r institution (including postal coa	ie and country)	
Manassas. Virginia United States of Ar	20110-2209		
Date of deposit M	arch 7, 1997	A	ccession Number 97924
C. ADDITIONAL	. INDICATIONS (leave blank i	f not applicable)	This information is continued on an additional sheet
D. DESIGNATED	STATES FOR WHICH IN	DICATIONS	ARE MADE (if the indications are not for all designated States)
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i crerence names.		1 *12 1111*	*** *	

(PCT Rule 13bis)

A. The indications made below relate to the micr on page 80	roorganism referred to in the description tine N/A
3. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet
Name of depositary institution American Ty	ype Culture Collection
Address of depositary institution (including postal 10801 University Boulevard Manassas, Virginia 20110-2209 United States of America	al code and country)
Date of deposit March 13, 1997	Accession Number 97958
C. ADDITIONAL INDICATIONS (leave b	blank if not applicable) This information is continued on an additional sheet
D. DESIGNATED STATES FOR WHIC	H INDICATIONS ARE MADE (if the indications are not for all designated States)
E. SEPARATE FURNISHING OF INDI	CATIONS (leave blank if not applicable)
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INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

(PCT Rule 13bis)

	ons made below relate to the m	nicroorganism referred in N/A	o in the description
3. IDENTIFIC	CATION OF DEPOSIT		Further deposits are identified on an additional sheet ()
Name of deposit	ary institution		1
•		Type Culture Collec	tion
Address of depo	sitary institution (including po	ostal code and country)	
10801 Univers	ity Boulevard ginia 20110-2209	•	
Date of deposit	May 22, 1997	A	ccession Number 209072
C. ADDITIO	NAL INDICATIONS (leave	e blank if noi applicable)	This information is continued on an additional sheet
D. DESIGNA	TED STATES FOR WHIC	CH INDICATIONS	ARE MADE (if the indications are not for all designated States)
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lame of depositary institution American Type Culture C	ollection
Address of depositary institution (<i>including postal code and coulogo</i> 10801 University Boulevard Manassas, Virginia 20110-2209 United States of America	intry)
Date of deposit September 4, 1997	Accession Number 209235
C. ADDITIONAL INDICATIONS (leave blank if not applic	cable) This information is continued on an additional sheet
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(PCT Rule 13bis)

A. The indications made below relate to the micro on page 84	organism referred to in the description
on page 64	Inc. 1474
B. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet
Name of depositary institution	
American 19	e Culture Collection
Address of depositary institution (including postal	code and country)
10801 University Boulevard Manassas, Virginia 20110-2209 United States of America	
Date of deposit August 28, 1997	Accession Number 209226
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INDICATIONS RELATING TO A DEPOSITED MICROORGANISM

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B. IDENTIFIC	ATION OF DEPOSIT		Further deposits are identified on an additional sheet
Name of deposita	ry institution America	in Type Culture Co	lection
10801 Universi	inia 20110-2209	postal code and coun	?ry)
Date of deposit	March 13, 1997		Accession Number 97957
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BNSDOCID: <WO 9842738A1>

Applicant's or agent's file reference number	Z004PCT	International application		
reference number		ا ق الحد	<u> </u>	

(PCT Rule 13bis)

A. The indications made below relate to the microorganism ref	ferred to in the description
B. IDENTIFICATION OF DEPOSIT	Further deposits are identified on an additional sheet []
Name of depositary institution American Type Culture C	Collection
Address of depositary institution (including postal code and co 10801 University Boulevard Manassas, Virginia 20110-2209 United States of America	puntry)
Date of deposit May 22, 1997	Accession Number 209073
C. ADDITIONAL INDICATIONS (leave blank if not appli	icable) This information is continued on an additional sheet
D. DESIGNATED STATES FOR WHICH INDICATI	IONS ARE MADE (if the indications are not for all designated States)
E. SEPARATE FURNISHING OF INDICATIONS de	ave blank if not applicable)
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Form PCT/XQ/134 (July 1992)

WO 98/42738 PCT/US98/05311

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What Is Claimed Is:

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- 1. An isolated nucleic acid molecule comprising a polynucleotide having a nucleotide sequence at least 95% identical to a sequence selected from the group consisting of:
- (a) a polynucleotide fragment of SEQ ID NO:X or a polynucleotide fragment of the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X;
- (b) a polynucleotide encoding a polypeptide fragment of SEQ ID NO:Y or a polypeptide fragment encoded by the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X;
- (c) a polynucleotide encoding a polypeptide domain of SEQ ID NO:Y or a polypeptide domain encoded by the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X;
- (d) a polynucleotide encoding a polypeptide epitope of SEQ ID NO:Y or a polypeptide epitope encoded by the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X;
- (e) a polynucleotide encoding a polypeptide of SEQ ID NO:Y or the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X, having biological activity;
 - (f) a polynucleotide which is a variant of SEQ ID NO:X;
 - (g) a polynucleotide which is an allelic variant of SEQ ID NO:X;
 - (h) a polynucleotide which encodes a species homologue of the SEQ ID NO:Y;
- (i) a polynucleotide capable of hybridizing under stringent conditions to any one of the polynucleotides specified in (a)-(h), wherein said polynucleotide does not hybridize under stringent conditions to a nucleic acid molecule having a nucleotide sequence of only A residues or of only T residues.
- 2. The isolated nucleic acid molecule of claim 1, wherein the polynucleotide fragment comprises a nucleotide sequence encoding a secreted protein.
- 3. The isolated nucleic acid molecule of claim 1, wherein the polynucleotide fragment comprises a nucleotide sequence encoding the sequence identified as SEQ ID NO:Y or the polypeptide encoded by the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X.

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4. The isolated nucleic acid molecule of claim 1, wherein the polynucleotide fragment comprises the entire nucleotide sequence of SEQ ID NO:X or the cDNA sequence included in ATCC Deposit No:Z, which is hybridizable to SEQ ID NO:X.

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5. The isolated nucleic acid molecule of claim 2, wherein the nucleotide sequence comprises sequential nucleotide deletions from either the C-terminus or the N-terminus.

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- 6. The isolated nucleic acid molecule of claim 3, wherein the nucleotide sequence comprises sequential nucleotide deletions from either the C-terminus or the N-terminus.
- 7. A recombinant vector comprising the isolated nucleic acid molecule of claim 1.
 - 8. A method of making a recombinant host cell comprising the isolated nucleic acid molecule of claim 1.
- 9. A recombinant host cell produced by the method of claim 8.
 - 10. The recombinant host cell of claim 9 comprising vector sequences.
- 11. An isolated polypeptide comprising an amino acid sequence at least 95% identical to a sequence selected from the group consisting of:
 - (a) a polypeptide fragment of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z;
 - (b) a polypeptide fragment of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z, having biological activity;

- (c) a polypeptide domain of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z;
- (d) a polypeptide epitope of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z;
- (e) a secreted form of SEQ ID NO:Y or the encoded sequence included inATCC Deposit No:Z;
 - (f) a full length protein of SEQ ID NO:Y or the encoded sequence included in ATCC Deposit No:Z;

- (g) a variant of SEQ ID NO:Y;
- (h) an allelic variant of SEQ ID NO:Y; or
- (i) a species homologue of the SEQ ID NO:Y.
- 12. The isolated polypeptide of claim 11, wherein the secreted form or the full length protein comprises sequential amino acid deletions from either the C-terminus or the N-terminus.
 - 13. An isolated antibody that binds specifically to the isolated polypeptide of claim 11.
- 14. A recombinant host cell that expresses the isolated polypeptide of claim
 11.
 - 15. A method of making an isolated polypeptide comprising:
- (a) culturing the recombinant host cell of claim 14 under conditions such that said polypeptide is expressed; and
 - (b) recovering said polypeptide.
 - 16. The polypeptide produced by claim 15.
 - 17. A method for preventing, treating, or ameliorating a medical condition, comprising administering to a mammalian subject a therapeutically effective amount of the polypeptide of claim 11 or the polynucleotide of claim 1.
- 25 18. A method of diagnosing a pathological condition or a susceptibility to a pathological condition in a subject comprising:
 - (a) determining the presence or absence of a mutation in the polynucleotide of claim 1; and
- (b) diagnosing a pathological condition or a susceptibility to a pathologicalcondition based on the presence or absence of said mutation.
 - 19. A method of diagnosing a pathological condition or a susceptibility to a pathological condition in a subject comprising:
 - (a) determining the presence or amount of expression of the polypeptide of claim 11 in a biological sample; and
 - (b) diagnosing a pathological condition or a susceptibility to a pathological condition based on the presence or amount of expression of the polypeptide.

- 20. A method for identifying a binding partner to the polypeptide of claim 11 comprising:
 - (a) contacting the polypeptide of claim 11 with a binding partner; and
- (b) determining whether the binding partner effects an activity of the polypeptide.
 - 21. The gene corresponding to the cDNA sequence of SEQ ID NO:Y.
- 10 22. A method of identifying an activity in a biological assay, wherein the method comprises:
 - (a) expressing SEQ ID NO:X in a cell;
 - (b) isolating the supernatant;
 - (c) detecting an activity in a biological assay; and
 - (d) identifying the protein in the supernatant having the activity.
 - 23. The product produced by the method of claim 22.

International Application No PCT/US 98/05311

CLASSIFICATION OF SUBJECT MATTER PC 6 C12N15/12 C12N5/10 C07K16/18 C07K14/47 ÎPC 6 C12N1/21 A61K38/17 G01N33/53 G01N33/50 G01N33/68 C12Q1/68 According to International Patent Classification (IPC) or to both national classification and IPC B. FIELDS SEARCHED Minimum documentation searched (classification system followed by classification symbols) CO7K C12Q GO1N A61K C12N Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practical, search terms used) C. DOCUMENTS CONSIDERED TO BE RELEVANT Relevant to claim No. Citation of document, with indication, where appropriate, of the relevant passages Category L. HILLIER ET AL.: "The WashU-Merck EST 1-3, X 7-11.21 project" EMBL SEQUENCE DATABASE, 2 July 1995, HEIDELBERG, FRG, XP002068365 y187a06.rl Homo sapiens cDNA clone 44938 5'; Accession no. H08241; L. HILLIER ET AL.: "The "WashU-Merck EST 1-3, X 7-11,21 project" EMBL SEQUENCE DATABASE, 26 August 1995, HEIDELBERG, FRG, XP002068366 ym94e01.rl Homo sapiens cDNA clone 166584 5', Accession no. R88485; 1-23 WO 97 07198 A (GENETICS INSTITUT) 27 Α February 1997 see the whole document -/--Further documents are listed in the continuation of box C. Patent family members are listed in annex. X Χ Special categories of cited documents: "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the "A" document defining the general state of the art which is not considered to be of particular relevance *E* earlier document but published on or after the international "X" document of particular relevance; the claimed invention filing date cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another "Y" document of particular relevance; the claimed invention citation or other special reason (as specified) cannot be considered to involve an inventive step when the document is combined with one or more other such docu document referring to an oral disclosure, use, exhibition or ments, such combination being obvious to a person skilled other means in the art document published prior to the international filing date but later than the priority date claimed "&" document member of the same patent family Date of the actual completion of the international search Date of mailing of the international search report 16, 09, 1998 17 June 1998 Authorized officer Name and mailing address of the ISA European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo ni, HORNIG H. Fax: (+31-70) 340-3016

International Application No
PCT/co 98/05311

		PCT/03 98	3/05311
C.(Continua	ation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Category °	Citation of document, with indication, where appropriate, of the relevant passages		Relevant to claim No.
A	WO 97 04097 A (GENETICS INST) 6 February 1997 see the whole document		1-23
A	US 5 536 637 A (JACOBS KENNETH) 16 July 1996 see the whole document	٠.	1-23
A	JACOBS K ET AL: "A novel method for isolating eukaryotic cDNA clones encoding secreted proteins." KEYSTONE SYMPOSIUM ON DENDRITIC CELLS: ANTIGEN PRESENTING CELLS OF T AND B LYMPHOCYTES, TAOS, NEW MEXICO, USA, MARCH 10-16, 1995. JOURNAL OF CELLULAR BIOCHEMISTRY SUPPLEMENT 0 (21A). 1995. 19. ISSN: 0733-1959, XP002027246 abstract no. C1-207 see abstract		1-23
A	WO 90 14432 A (GENETICS INST) 29 November 1990 see the whole document		1-23
Α	WO 96 17925 A (IMMUNEX CORP) 13 June 1996 see the whole document		1-23
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Inte .ional application No.

PCT/US 98/05311

Box i	Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This Inte	ernational Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. X	Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely: Remark: Although claim 17 is directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
2.	Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3.	Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box II	Observations where unity of invention is lacking (Continuation of item 2 of first sheet)
This Inte	ernational Searching Authority found multiple inventions in this international application, as follows:
se	ee further information sheet
1.	As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2.	As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3.	As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. X	No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
	see further information sheet
Remai	rk on Protest The additional search fees were accompanied by the applicant's protest.
	No protest accompanied the payment of additional search fees.

Form PCT/ISA/210 (continuation of first sheet (1)) (July 1992)

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

Invention 1. Claims: (1-23) partially

-An isolated nucleic acid molecule comprising a polynucleotide having a nucleotide sequence at least 95% identical to a sequence selected from SEQ ID no. 11; wherein said polynucleotide fragment comprises a nucleotide sequence encoding a secreted protein encoding the sequence SEQ ID no.125 or the polypeptide encoded by the cDNA sequence included in ATCC Deposit nos: 97923/209071, which is hybridizable to SEQ ID no.11; a recombinant vector comprising said isolated nucleic acid molecule; a method of making a recombinant host cell comprising said isolated nucleic acid molecule; a recombinant host cell comprising said vector; an isolated polypeptide comprising an amino acid sequence at least 95% identical to a sequence SEQ ID no. 125; an isolated antibody that binds specifically to said isolated polyeptitde; a recombinant host cell that expresses said isolated polypeptide; a method of making said polypeptide; a method for preventing, treating or ameliorating a medical condition which comprises administering to a mammalian subject a therapeutically effective amount of said polypeptide; a method of diagnosing a pathological condition or a susceptibility to a pathological condition in a subject using said polynucleotide and/or polypeptide sequence; a method for identifying a binding partner to said polypeptide; a gene corresponding to the cDNA sequence of SEQ ID no.11; a method for identifying an activity in a biological assay, by using the expression of SEQ ID no. 125;

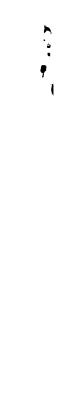
Inventions 2 to 87. Claims: (12-23) partially

-Idem as subject 1 but limited to gene nos. 2 to 87 respectively cDNA clone sequences HAGFY16/HBMCF37/HFLQB16 to HCED021. (Invention 2 is limited to SEQ ID nos. 12,98,99,126,212 and 213; Invention 3 is limited to SEQ ID nos.13 and 127;; Invention 87 is limited to SEQ ID nos.97 and 211;)

Int ation on patent family members

International Application No
PCT/US 98/05311

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
WO 9707198 A	27-02-97	US 5707829 A AU 6712396 A AU 6768596 A EP 0839196 A EP 0851875 A WO 9704097 A	18-02-97 12-03-97 06-05-98 08-07-98
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